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(54) Resurfacing of rodent antibodies.

- (57) A method for determining how to humanize a rodent antibody or fragment thereof by resurfacing, said method comprising:
 - (a) determining the conformational structure of the variable region of said rodent antibody or fragment thereof by constructing a three-dimensional model of said rodent antibody variable region;
 - (b) generating sequence alignments from relative accessibility distributions from x-ray crystal-lographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of heavy and light chain framework positions wherein said set is identical in 98% of said sufficient number of rodent antibody heavy and light chains;
 - (c) defining for said rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using said set of framework positions generated in said step (b);
 - (d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c), wherein said heavy and light chain from said human antibody are or are not naturally paired;
 - (e) substituting, in the amino acid sequence of said rodent antibody or fragment thereof to be humanized said set of heavy and light chain surface exposed amino acid residues defined in said step (c) with said set of heavy and light chain surface exposed amino acid residues identified in said step (d);
 - (f) constructing a three-dimensional model of said variable region of said rodent antibody or fragment thereof resulting from the substituting specified in said step (e):
 - (g) identifying, by comparing said three-dimensional models constructed in said steps (a) and (f), any amino acid residues from said set identified in said step (d), that are within 5 Angstroms of any atom of any residue of the complementarity determining regions of said rodent antibody or fragment thereof to be humanized; and
 - (h) changing any residues identified in said step (g) from the human to the original rodent amino acid residue to thereby define a rodent antibody humanizing set of surface exposed amino acid residues; with the proviso that said step (a) need not be conducted first, but must be conducted prior to said step (g).

FIELD OF THE INVENTION

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The present invention relates to the development of prediction rules that can be used to accurately model the variable regions (V-regions) of antibodies. The development of these rules and their application in the predictive molecular restructuring of the surfaces of variable domains of non-human monoclonal antibodies enables changing of the surface, resurfacing, of these monoclonal antibody V-regions to replicate the surface characteristics found on human antibody V-regions. This method of resurfacing non-human monoclonal antibody V-regions to resemble human antibody V-regions is expected to permit the production of functional altered antibodies, which retain the binding parameters of the original non-human monoclonal antibody, with improved therapeutic efficacy in patients due to the presentation of a human surface on the V-region.

BACKGROUND OF THE INVENTION

General Background of Antibodies

Murine monoclonal antibodies are widely used as diagnostic and therapeutic agents in the treatment of human disease. Mice can be readily immunized with foreign antigens to produce a broad spectrum of high affinity antibodies. Invariably, the introduction of murine or other rodent antibodies into humans results in the production of a human anti-mouse antibody (HAMA) response due to the presentation of a foreign protein in the body. The production of HAMA in patients can result from the introduction of foreign antibody in a single dose or from extended use in therapy, for example, for the treatment of cancer. Extended use of murine antibody is generally limited to a term of days or weeks in patients before concerns of anaphylaxis arise. Moreover, once HAMA has developed in a patient, future use of murine antibodies for diagnostic or therapeutic purposes is often precluded for the same reasons.

Beyond ethical considerations, attempts to produce human monoclonal antibodies have not been highly successful for a number of reasons. The production *in vitro* of human monoclonals rarely results in high affinity antibodies. *In vitro* cultures of human lymphocytes yield a restricted range of antibody responses relative to the broad spectrum of reactive antibodies produced *in vivo* through direct immunization of mice. Additionally, in humans, immune tolerance prevents the successful generation of antibodies to self-antigens. All of these factors have contributed to the search for ways to modify the structures of murine monoclonal antibodies to improve their use in patients. Many investigators have attempted to alter, reshape or humanize murine monoclonal antibodies in an effort to improve the therapeutic application of these molecules in patients.

Strategies of Antibody Humanization

The earliest reports of the controlled rearrangement of antibody domains to create novel proteins was demonstrated using rabbit and human antibodies as described by Bobrzecka, K. et al. (Bobrzecka, K., Konieczny, L., Laidler, P. and Rybarska, J. (1980), Immunology Letters 2, pp. 151-155) and by Konieczny et al. (Konieczny, L., Bobrzecka, K., Laidler, P. and Rybarska, J. (1981), Haematologia 14 (I), pp. 95-99). In those reports, the protein subunits of antibodies, rabbit Fab fragments and human Fc fragments, were joined through protein disulfide bonds to form new, artificial protein molecules or chimeric antibodies.

Recombinant DNA technology was used to construct gene fusions between DNA sequences encoding mouse antibody variable light and heavy chain domains and human antibody light chain (LC) and heavy chain (HC) constant domains to permit expression of the first recombinant "near-human" antibody (chimeric antibody) product (Morrison, S.L., Johnson, M.J., Herzenberg, L.A. and Oi, V.T. (1984), Proc. Natl. Acad. Sci. U.S.A. 81, pp. 6851-6855).

The kinetics and immune response in man to chimeric antibodies has been examined (LoBuglio, A.F., Wheeler, R.H., Trang, J., Haynes, A., Rogers, K., Harvey, E.B., Sun, L., Ghrayeb, J. and Khazaeli, M.B. (1989), Proc. Natl. Acad. Sci. **86**, pp. 4220-4224).

Chimeric antibodies contain a large number of non-human amino acid sequences and are immunogenic in man. The result is the production of human anti-chimera antibodies (HACA) in patients. HACA is directed against the murine V-region and can also be directed against the novel V-region/C-region (constant region) junctions present in recombinant chimeric antibodies.

To overcome some of the limitations presented by the immunogenicity of chimeric antibodies, the DNA sequences encoding the antigen binding portions or complementarity determining regions (CDR's) of murine monoclonal antibodies have been grafted by molecular means in the DNA sequences encoding the frameworks of human antibody heavy and light chains (Jones, P.T., Dear, P.H., Foote, J., Neuberger, M.S. and Winter, G. (1986), Nature 321, pp. 522-525; Riechmann, L., Clark, M., Waldmann, H. and Winter, G. (1988), Nature 332,

pp. 323-327). The expressed recombinant products called reshaped or humanized antibodies are comprised of the framework of a human antibody light or heavy chain and the antigen recognition portions, CDR's, of a murine monoclonal antibody. Several patent applications have been filed in this area including, for example, European Patent Application, Publication No. 0239400; European Patent Application, Publication Nos. 0438310A1 and 0438310A2; International Patent Publication No. WO 91/09967; and International Patent Publication No. WO 90/07861.

However, it is questionable whether European Patent Application (EP), Publication No. 0239400 is truly enabling. It is not assured in this patent that the best fit is made to assure proper presentation of the CDR loops at the antibody combining site.

EP Publication Nos. 0438310A1 and 0438310A2 go a step beyond EP Publication No. 0239400 by protecting the importance of uniquely selected human frameworks for the human light chain (LC) and heavy chain (HC) V-regions. These V-region frameworks should show a high degree of sequence similarity with the frameworks of the murine monoclonal antibody and present the CDR's in the appropriate configuration. However, the criteria for sequence matching are no more sophisticated than simple homology searching of the antibody protein or DNA databases.

International Patent Publication No. WO 91/09967 attempts a further variation of the method disclosed in EP Publication No. 0239400. In International Patent Publication No. WO 91/09967, homology of the donor sequences and the acceptor framework is not important, rather it discloses that a selected set of residues in the LC and HC are critically important to humanization. The ability to make changes at these positions is the basis of International Patent Publication No. WO 91/09967.

International Patent Publication No. WO 90/07861 proposes four important criteria for designing humanized antibodies. 1) Homology between human acceptor and non-human donor sequences. 2) Use donor rather than acceptor amino acids where the acceptor amino acid is unusual at that position. 3) Use donor framework amino acids at positions adjacent to the CDR. 4) Use donor amino acids at framework positions where the sidechain atom is within 3 Angstroms of the CDR in a 3-D model. The first antibody humanized by this method retained less than 1/3 the affinity of the original monoclonal antibody.

None of the above methods for designing a humanized antibody are predictable due to the questions that surround CDR framework interactions. By replacement of murine framework with human framework, there is no guarantee of identical conformations for CDR's because i) the V_L - V_H interaction is not identical in all V-regions and ii) accurate prediction of the CDR-framework interactions are key to faithful reproduction of the antigen binding contacts.

The above methods do not offer a general solution to solving the issues surrounding antibody humanization, rather the methods as outlined in each reference above involve a substantial amount of trial and error searching to obtain the desired affinity in the final humanized product. More importantly, there is no guarantee that corrective changes in framework amino acids will leave the reshaped V-regions resembling the surface character of a truly human antibody. Therefore, it can be argued that antibodies humanized by the above methods may be immunogenic in man.

Antigenicity of Antibodies

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The antigenicity/immunogenicity of an antibody, including recombinant reshaped antibody products, introduced into humans can be viewed as a surface phenomenon. In general one can view the immune system as scanning the surface of a protein introduced to the body. If the F_V portion of a humanized antibody 'opensup' in the circulation then internal residues can be presented to the immune system. On the other hand, if the F_V portion is stable and tightly packed then only the surface residues presented by the V-regions and the interface between the V_L and V_H regions will be 'scanned'.

Surface Reshaping or Resurfacing of Antibobies

The notion of surface presentation of proteins to the immune system raises the prospect of redesigning murine monoclonal antibodies to resemble human antibodies by humanizing only those amino acids that are accessible at the surface of the V-regions of the recombinant F_V. The resurfacing of murine monoclonal antibodies to reduce their immunogenicity could be beneficial in maintaining the avidity of the original monoclonal antibody in the reshaped version, because the natural framework-CDR interactions are retained. The value of maintaining the integrity of the framework-CDR interactions has been illustrated as summarized below.

In a recent research report, two different reshaped versions of the rat monoclonal antibody, Campath-9 (anti-human CD4), were generated (Gorman, S.D., Clark, M.R., Routledge, E.G., Cobbold, S.P. and Waldmann, H. (1991), Proc. Natl. Acad. Sci. U.S.A. 88, pp. 4181-4185). In one version, pV_HNEW/C_{G1}, the acceptor V_H fra-

mework was from the human NEW-based heavy chain, which has 47% identical residues to the Campath-9 V_H . While in the second version, pV_HKOL/C_{G1} , the acceptor V_H framework was from the human KOL antibody, which has 72% identical residues to Campath-9 V_H . Each reshaped antibody contained the identical V_L domain from the human REI antibody sequence. However, the recombinant product of pV_HKOL/C_{G1} had an avidity for CD4 that was substantially greater than the product of pV_HNEW/C_{G1} . The authors proposed a reshaping strategy where human sequences, that are highly homologous to the rodent antibody of interest, are transferred, by in vitro mutagenesis, into the rodent V-region to create a "bestfit" reshaped antibody. This strategy uses the term "bestfit" to describe the modeling process, however, there is no quantitative formula employed to assess "bestfit", and so in effect, the process is subjective. Additionally, there is no resurfacing concept presented in that paper.

The concept of reducing rodent-derived antibody immunogenicity through the replacement of exposed residues in the antibody framework regions which differ from those of human origin is discussed in a recent paper (Padlan, E.A. (1991), Molecular Immunology 28, pp. 489-498). In that paper, the variable domains of two antibody structures, KOL (human) and J539 (mouse), are examined. The crystal structures of the Fab fragments of these two antibodies have been elucidated to high resolution. The solvent accessibility of the exposed framework residues in the variable domains of these two antibodies were compared to a sequence database of human and murine antibody V-region subgroups. On the basis of his findings, Padlan proposed to reduce the antigenicity of allogeneic variable domains [murine V-regions], through replacement of the exposed residues in the framework regions with residues usually found in human antibodies. In murine sequences with the highest similarity to a given human sequence, the number of changes necessary to "humanize" a murine V-region surface would range from 6-15 amino acid changes per V-region. This reference suggests how to convert one antibody surface into another but no general method is developed. Application of the procedure is provided by two examples, a worst-case and a best-case.

25 Worst Case:

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Among the representative murine kappa V_L sequences examined for which its autologous V_H has been sequenced, $S107V_L$ has the most residues that need to be replaced to humanize it. $S107V_L$ is most similar to the members of the human subgroup VKIV and JK2. The exposed or partially exposed residues that need to be replaced are those at positions $\underline{9}$, 10, 14, $\underline{15}$, 16, 17, 18, $\underline{22}$, $\underline{41}$, $\underline{63}$, $\underline{80}$, $\underline{85}$, 100 and 106. Murine V-region $S107V_H$ is most similar in its framework to the members of the human subgroup VHIII and JH6. The exposed or partially exposed residues in $S107V_H$ that need to be replaced are those at positions 3, 40, 68, 73, 75, 76, 82b and 89. A total of 23 residues need to be replaced to humanize the variable domains of S107.

Best Case:

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Among the murine V_H sequences examined for which the autologous V_L has also been sequenced, MOPC21 V_H has the least number of residues that need to be replaced to humanize it. MOPC21 V_H is most similar in its framework to the members of the human subgroup HIII and JH6. The exposed or partially exposed residues that need to be replaced are those at positions 1, 42, 74, 82a, 84, 89 and 108. MOPC21 V_L is most similar in its framework to human subgroup VKIV and JK4. The exposed or partially exposed residues that need to be replaced are those at positions 1, 9, 12, 15, 22, 41, 63, 68, 83 and 85. A total of 17 amino acids need to be replaced to humanize the variable domains of MOPC21.

Of the light chains in the Best- and Worst-Case examples cited above, $S107V_L$ required changes at 15 positions and $MOPC21V_L$ required changes at 10 positions. Only seven of the changes are common to both of these light chain sequences (see underlined residues). Moreover, of the heavy chain residues that need to be replaced to humanize the respective V-regions, $S107V_H$ required changes at 8 positions and $MOPC21V_H$ required changes at 7 positions. In this instance, only one position is common to both of these heavy chain sequences (see residues in boldface).

An analysis of S107 V-regions alone would not have led to the prediction of which residues to change in MOPC21. The reason for this is that the surface residues in Padlan's analysis are only determined by reference to the crystal structure analysis of <u>one</u> antibody. In addition, the basis for defining the surface exposure of an amino acid at a particular position on that crystal structure is a continuous gradient of change, e.g., the fractional solvent accessibility values (Padlan, E.A. (1990), Molecular Immunology 28, pp. 489-498) were computed, where: 0 to 0.2 = completely buried, 0.2 to 0.4 = mostly buried, 0.4 to 0.6 = partly buried/partly exposed, 0.6 to 0.8 = mostly exposed, and 0.8 or above = completely exposed. By limiting the analysis of exposed surface residues to a single crystal structure and by superimposing a broad range of solvent accessibility ratios on exposed residues, such a modeling strategy could be expected to have a wide margin of error in its calculations.

This model fails to take into account the great majority of structural information available in the database for other antibody crystal structures.

SUMMARY OF THE INVENTION

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Accordingly, it is an object of this invention to provide humanized rodent antibodies or fragments thereof, and in particular, humanized rodent monoclonal antibodies that have improved therapeutic efficacy in patients due to the presentation of a human surface on the V-region. This and other objects have been attained by providing a method for determining how to humanize a rodent antibody or fragment thereof by resurfacing the method comprising:

- (a) determining the conformational structure of the variable region of the rodent antibody or fragment thereof by constructing a three-dimensional model of the rodent antibody variable region;
- (b) generating sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of heavy and light chain framework positions wherein the set is identical in 98% of the sufficient number of rodent antibody heavy and light chains;
- (c) defining for the rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using the set of framework positions generated in step (b);
- (d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to the set of surface exposed amino acid residues defined in step (c), wherein the heavy and light chain from the human antibody are or are not naturally paired;
- (e) substituting, in the amino acid sequence of the rodent antibody or fragment thereof to be humanized the set of heavy and light chain surface exposed amino acid residues defined in step (c) with the set of heavy and light chain surface exposed amino acid residues identified in step (d);
- (f) constructing a three-dimensional model of the variable region of the rodent antibody or fragment thereof resulting from the substituting specified in step (e);
- (g) identifying, by comparing the three-dimensional models constructed in steps (a) and (f), any amino acid residues from the set identified in step (d), that are within 5 Angstroms of any atom of any residue of the complementarity determining regions of the rodent antibody or fragment thereof to be humanized; and
- (h) changing any residues identified in step (g) from the human to the original rodent amino acid residue to thereby define a rodent antibody humanizing set of surface exposed amino acid residues; with the proviso that step (a) need not be conducted first, but must be conducted prior to step (g).

Also provided is a method for producing a humanized rodent antibody or fragment thereof from a rodent antibody or fragment thereof, the method comprising:

- (I) carrying out the above-described method for determining how to humanize a rodent antibody or fragment thereof by resurfacing; and
- (II) modifying the rodent antibody or fragment thereof by replacing the set of rodent antibody surface exposed amino acid residues with the rodent antibody humanizing set of surface exposed amino acid residues defined in step (h) of the above-described method.
- In a preferred embodiment, the rodent antibody or fragment thereof is a murine antibody, and most preferably murine antibody N901.

BRIEF DESCRIPTION OF THE FIGURES

Figure 1 shows an algorithm that can be used for constructing a three-dimensional model of the rodent antibody variable region.

Figure 2 is a diagram showing the approach to determine how to humanize a rodent antibody or fragment thereof according to the present invention.

Figures 3A and 3B are plots of relative accessibility of amino acid residues for twelve antibody F_V structures, mapped onto the sequence alignment of these structures. Structures Glb2 (Jeffrey, P.D., Doctor of Philosophy Thesis, University of Oxford, United Kingdom, 1991), D1.3 (Amit, A.G., Mariuzza, R.A., Phillips, S.E.V. and Poljak, R.J. (1986), Science 233, pp. 747-753), 3D6 (Grunow, R., Jahn, S., Porstman, T., Kiessig, T., Steinkeller, H., Steindl, F., Mattanovich, D., Gurtler, L., Deinhardt, F., Katinger, H. and von R., B. (1988), J. Immunol. Meth. 106, pp. 257-265) and 36-71 (5fab) (Rose, D.R., Strong, R.K., Margolis, M.N., Gefter, M.L. and Petsko, G.A. (1990), Proc. Natl. Acad. Sci. U.S.A. 87, pp. 338-342) are not yet present in the Brookhaven database. The other structures used were: 2hfl (Sheriff, S., Silverton, E.W., Padlan, E.A., Cohen, G.H., Smith-Gill, S.J., Finzel, B.C. and Davies, D.R. (1987), Proc. Natl. Acad. Sci. U.S.A. 84, pp. 8075-8079), 3hfm (Padlan, E., Silverton, E., Sheriff, S., Cohen, G., Smith-Gill, S. and Davies, D. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, pp.

5938-5942), 2fbj (Mainhart, C.R., Potter, M. and Feldmann, R.J. (1984), Mol. Immunol. 21, pp. 469-478), 3fab (Saul, F.A., Amzel, L.M. and Poljak, R.J. (1978), J. Biol. Chem. 253, pp. 585-597), 4fab (Herron, J., He, X., Mason, M., Voss, E. and Edmunson, A. (1989), Proteins: Struct., Funct., Genet. 5, pp. 271-280), 2mcp (Segal, D., Padlan, E., Cohen, G., Rudikoff, S., Potter, M. and Davies, D. (1974), Proc. Natl. Acad. Sci. U.S.A. 71, pp. 4298-[??]), 2fb4 (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391), and 1f19 (Lascombe, M. Alzari, P., Boulot, G., Salujian, P., Tougard, P., Berek, C., Haba, S., Rosen, E., Nisonof, A. and Poljak, R. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, p. 607). These structures are designated by their Brookhaven entry code. The sequence numbering used here is described in Figures 4A and 4B. Figure 3A graphically shows the relative accessibility for the heavy chain and Figure 3B graphically shows the relative accessibility for the light chain.

Figures 4A and 4B show alignments of sequences generated using the three methods of humanization. Sequences are: 1) Original rodent N901. 2+3) KOL (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391) and reshaped N901 using KOL surface. 4+5) Most homologous sequences, L(KV2F) (Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res. pp. 6499-6513) and H(G36005) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87), and reshaped N901 using these sequences. 6+7) Most homologous with respect to surface residues, L(KV4B) (Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. 3, pp. 6515-6529) and H(PLO123) (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. 168, pp. 229-245), and reshaped N901 using these sequences. The numbering is the same as used in the antibody modelling program ABM (trademark for commercial software, Oxford Molecular Ltd., Oxford, U.K.), which is based on structural conservation and not sequence homology as used by Padlan et al. (Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition). The sequence changes which have to be introduced in order to resurface N901 with a given sequence are marked with bars, back-mutations as determined from F_V models are marked with stars. The sequence homology of given sequences to N901 are shown in brackets after each sequence

Figure 5 is a stereo plot of mean antibody β -barrel, coordinates determined by iterative multiple fitting of eight antibody structures. Strands 7 and 8 comprise the 'take off' positions for CDR H3 and are not included in the fitting of V_L and V_H regions.

Figure 6 is a plot of RMS deviation from the mean of the eight β -sheet strands comprising the framework. The RMS was calculated from structures F19.9, 4-4-20, NEW, FBJ, KOL, HyHEL-5, HyHEL-10 and McPC603. N,C α ,C atoms are included in the plot. The residues used are shown in the alignment (Table 2). The most disordered residues are all the residues of strand HFR4, the last residue of LFR1, and the first and last residue of HFR2. The nomenclature of the strands is explained in the alignment in Table 2. LFR1 - #1, LFR2 - #2, LFR3 - #3, LFR4 - #4, HFR1 - #5, HFR2 - #6, HFR3 - #7, HFRS4 - #8.

Figure 7 is a flowchart of the overall modelling protocol known as CAMAL.

Figure 8 is a plot of superimposed loop backbones for models and x-ray structures discussed in Example 2. The loops are positioned after global framework fit. This does not represent the best local least squares fit, but shows how the loops are positioned globally onto the framework.

Figures 9A to 9D are stereo (N,C- α ,C,O) representations of crystal structures and models of D1.3, 3671 and Gloop-2 variable domain and β -barrel strands described in Example 2. Crystal structures are shown with open bonds, model with solid bonds. The difference between the 3D6-H3 in the model and the crystal structure is due to a 5-7° twist in the extended β -sheet conformation of this loop, Figure 9A: D1.3, Figure 9B: 36-71, Figure 9C: Gloop-2, Figure 9D: 3D6.

Figure 10 is a histogram showing the distribution of loop length for CDR H3 loops, data from Kabat et al. (Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition).

DETAILED DESCRIPTION OF THE INVENTION

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The existence of specific, yet different, surface patches in murine and human antibodies may be the origin of the inherited immunogenicity of murine antibodies in humans. Statistical analysis of a database of unique human and murine antibody F_V fragments has revealed that certain combinations of residues in exposed surface positions are specific for human and murine sequences. The combinations are not the same in human and murine F_V domains. However, it is possible to define families of surface residues for the two species of antibodies. These families reveal a novel method for the "humanization" or reshaping of murine antibodies. Humanization is the modification of the solvent accessible surface of a non-human antibody or fragment thereof to resemble the surface of a chosen human antibody or fragment thereof such that the modified non-human antibody or fragment thereof exhibits lower immunogenicity when administered to humans. Such a process

applies in the present application to antibody variable regions but could equally well apply to any other antibody fragment. The method is considered to be generally applicable to humanization of rodent antibodies.

According to the present invention, a statistical analysis is presented which is based on accessibility calculated for a range of antibody crystal structures. When this information is applied to an antibody sequence database, it is possible to discriminate between human and murine antibodies at the sequence level purely on the basis of their surface residue profiles.

Rational Resurfacing Approach

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There are several key features of the resurfacing approach of the present invention.

- 1) This method uses as a starting point, construction of a three-dimensional model of a rodent variable region by known methods;
- 2) A large number (e.g., twelve) of antibody F_V or Fab fragment x-ray crystallographic structures are analyzed to produce an unambiguous set of surface exposed amino acid residues that will be positionally identical for a majority (98%) of antibodies. The set is produced by identifying all those residues whose solvent accessibility is above a given cut-off (typically 30%), calculated using a modification of the method of Kabsch and Sander (Kabsch, W. and Sander, C. (1983), Biopolymers 22, pp. 2257-2637) in which explicit atomic radii are used for each atom type to predict sidechain positions as is described below in more detail; 3) Using a complete human antibody database, the best set of human heavy and light chain surface exposed amino acid residues is selected on the basis of their closest identity to the set of surface amino acid residues of the murine antibody:
- 4) In order to retain the conformational structure- of the CDRs of the rodent antibody, replacement of any human surface exposed amino acid with the original rodent surface exposed amino acid residue is carried out whenever a surface residue is calculated from the three-dimensional model to be within 5 Angstroms of a CDR residues.

The general resurfacing approach of the present invention is illustrated in Figure 2. The approach can be divided into two stages. In the first, the rodent framework (white) is retained and only the surface residues changed from rodent (dark grey circles) to the closest human pattern (light grey circles). This should remove the antigenicity of the rodent antibody. In the second stage, surface residues within 5 Angstroms of the CDRs are replaced with the rodent equivalents in an attempt to retain antigen binding and CDR conformation.

The method of the present invention is applicable to whole antibodies as well as antibody fragments. Suitable antibody fragments that can be used can readily be determined by the skilled artisan. Examples of some suitable fragments include a single chain antibody (SCA), an antibody F_V fragment, Fab fragment, Fab₂ fragment, Fab' fragment, or other portion of an antibody comprising the binding site thereof.

According to the present invention, an important step in the method for determining how to modify a rodent antibody or fragment thereof by resurfacing is to determine the conformational structure of the variable region of the rodent antibody or fragment thereof to be humanized by constructing a three-dimensional model of the rodent antibody variable region. This can be done by known methods such as those described, for example, in Martin et al. (Martin, A.C.R., Cheetham, J.C. and Rees, A.R. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, pp. 9268-9272; Methods in Enzymology (1991), 203, pp. 121-152) and as described in detail in Example 2.

Martin et al. describe an algorithm which is depicted in Figure 1. The algorithm applies to murine and human antibodies equally well. The present inventors therefore expect that, based on sequence similarity between antibodies of different species (Kabat, E.A. Segments of Proteins of Immunological Interest, National Institutes of Health, U.S.A. 1991), the algorithm will work equally well for rat and other rodent antibodies.

Briefly, the algorithm depicted in Figure 1 can be summarized as follows. The framework region of an antibody to be modelled is selected on the basis of sequence homology and constructed by a least squares fit onto the six conserved strands of the variable region β -barrel. Light and heavy chain complementarity determining regions are constructed using a combination of canonical structures (Chothia, C. and Lesk, A.M. (1987), J. Molec. Bio. **196**, pp. 901-917), database searching and conformational searching. Detailed descriptions of these methods are described in Example 2 herein and in the above two references (Martin et al. 1989 and 1991).

According to the present invention, another three-dimensional model is also constructed. The other three-dimensional model is of the rodent antibody variable region having human antibody surface amino acid residues substituted therein at particular rodent antibody surface residue positions.

This other three-dimensional model is constructed by carrying out the series of steps described next.

The first of the steps is to generate sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of framework positions of surface exposed amino acid residues which is identical in a majority

(98%) of the variable regions.

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As used herein, the term "framework" means the antibody variable region from which the complementarity determining regions have been excluded.

"Complementarity determining regions" means those amino acid sequences corresponding to the following numbering system as defined by Kabat, E.A. (In Sequences of Immunological Interest, N.I.H., U.S.A., 1991).

· · · · · · · · · · · · · · · · · · ·			
Light Chain	L1	residues	24-34
Light Chain	L2	residues	50-56
Light Chain	L3	residues	89-97
Heavy Chain	H1	residues	31-358
Heavy Chain	H2	residues	50-58
Heavy Chain	Н3	residues	95-102

A sufficient number of rodent antibody fragments that need to be analyzed in order to produce the set of framework positions of surface exposed amino acid residues can readily be determined by the skilled artisan through routine experimentation using a database of antibody sequences. Thus, this step can be conducted using suitable databases now in existence or later compiled.

The x-ray crystallographic structures are used to determine relative accessibility distributions of surface exposed amino acid residues. The relative accessibility distributions identify all those residues whose solvent accessibility is above a given cut-off (typically 30%), calculated using a modification of the method of Kabsch and Sander (Kabsch, W. and Sander C. (1983), Biopolymers 22, pp. 2257-2637) in which explicit atomic radii are used for each atom type.

The relative accessibility distributions determined from the x-ray crystallographic structures can then be used to generate sequence alignments which give a set of framework positions of surface exposed amino acid residues which is identical in a majority (98%) of the variable regions.

The set of framework positions of surface exposed amino acid residues for the variable regions of murine antibodies is shown in Table 1, set forth in Example 1, and was produced using the sequence alignments and accessibility distributions shown in Figures 3A and 3B.

Once a set of framework positions of surface exposed amino acid residues for the variable regions of the rodent antibodies have been generated, the surface exposed residues of the heavy and light chain pair of the rodent antibody, or fragment thereof, to be humanized can be identified using an alignment procedure such as that described in Example 1 and shown in Figures 3A and 3B. This defines a set of surface exposed amino acid residues of a heavy and light chain pair of a rodent antibody or antibody fragment to be humanized.

Next, a complete human antibody sequence database is used to identify a set of surface exposed amino acid residues from a human antibody variable region that have the closest positional identity to the set of surface exposed amino acid residues of the variable region of the rodent antibody that is to be humanized. The set of surface exposed amino acid residues from the human antibodies can be separately identified for a heavy chain and for a light chain that are not naturally paired and/or a set can be identified from a natural human heavy and light chain pair, that is, a pair originating from a single B cell or hybridoma clone. Preferably, the set is one from a natural human heavy and light chain pair.

A humanized rodent antibody that gives the appearance of a human antibody is then predicted by substituting the set of surface exposed amino acid residues from the rodent antibody or fragment thereof to be humanized with the set of surface exposed amino acid residues from the human antibody.

A three-dimensional model can then be constructed from the resulting, fully substituted variable region of the rodent antibody or fragment thereof. The three-dimensional model is constructed using the same known methods mentioned above for constructing a 3-D model of the original rodent antibody or fragment thereof.

While the antigenicity of this fully "resurfaced" or humanized antibody should be removed, an additional factor to be addressed is the binding affinity or the binding strength of the resurfaced antibody. Changes in the framework of the variable domain introduced through resurfacing can influence the conformation of the CDR loops and therefore antigen binding of the antibody. According to the present invention, this problem is removed by the next step which is to identify, by means of a comparison of both of the above-described threedimensional models of the rodent antibody variable region, any residues from the set of surface exposed amino acid residues of the variable region heavy and light chain pair of the human antibody identified that are within 5 Angstroms of any atom of any residue of the rodent antibody or antibody fragment complementarity determining regions (CDRs).

Any residue(s) so identified is then changed back from the human to the original rodent amino acid residue(s).

The results of this method can then be applied to a particular rodent antibody by well known methods. Briefly, genes for the humanized variable heavy and light chain regions are constructed using standard recombinant DNA methods (Sambrook, J., Fritsch, E.F. and Maniatis, T. (1989), Molecular Cloning, Second Edition). For example, a PCR method can be used (Daugherty et al. (1991), Nucleic Acids Research 19, pp. 2471-2476).

Variable heavy chain or variable light chain gene constructs are subcloned into appropriate expression vectors. Suitable expression vectors contain either a human gamma or human kappa constant region gene, a suitable promoter, a sequence coding for a human immunoglobulin leader peptide (for example: met-gly-trp-ser-cys-ile-ile-leu-phe-leu-val-ala-thr-ala-thr (SEQ ID NO:39), Olandi et al. (1989), PNAS 86, pp. 3833-3837), and a drug selectable marker.

Heavy and light chain expression plasmids can be co-transfected, for example, by electroporation into suitable cells, for example, SP2/0 cells, and selected with an appropriate drug, G418, for example. Screening for intact antibody can be accomplished by ELISA assay. 96-well plates are coated with, for example, goat antihuman kappa chain antibody, and light chains are detected with, for example, goat anti-human antibody conjugated to alkaline phosphatase.

As another approach, light chain constructs are transfected, for example, by electroporation into suitable cells, for example, SP2/0 cells and selected, for example, in hygromycin. Screening for light chain expression can be accomplished by ELISA assay. 96-well plates are coated with, for example, goat anti-human kappa chain antibody, and light chains are detected with, for example, goat anti-human antibody conjugated to alkaline phosphatase.

A light chain producing line is then used as a host to electroporate in the heavy chain construct. The heavy chain plasmid is co-transfected with a plasmid containing the gene coding for another drug marker, for example, neomycin resistance and selected in the presence of the drug G418. Screening for intact antibody is accomplished by ELISA assay. 96-well plates are coated with, for example, goat anti-human Fc and detected with, for example, goat anti-human light chain conjugated to alkaline phosphatase.

EXAMPLE 1 AND COMPARATIVE EXAMPLES

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The superiority of the presently claimed method for determining how to modify a rodent antibody or fragment thereof by resurfacing in order to produce a humanized rodent antibody will now be described by reference to the following example and comparative examples which are illustrative and are not meant to limit the present invention.

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A) Analysis for Murine Antibodies

In order to determine the positions which are usually accessible on the surface of the F_V domain of murine antibodies, the accessibility was calculated for twelve Fab x-ray crystallographic structures obtained from the Brookhaven database (Bernstein, F., Koetzle, T., Williams, G., Meyer, E., Brice, M., Rodgers, J., Kennard, O., Shimanouchi, T. and Tasumi, M. (1977), J. Mol. Biol. 112, pp. 535-542). The relative accessibility was calculated using the program MC (Pedersen, J. (1991)), which implements a modified version of the DSSP (Kabsch, W. and Sander, C. (1983), Biopolymers 22, pp. 2257-2637) accessibility calculation routine in which explicit atomic radii are specified for every atom. A residue was defined as being surface accessible when the relative accessibility was greater than 30%. The alignment positions of these residues were conserved in all twelve structures (98% identity). Surface accessible framework positions constitute 40% of the F_V surface area. The remaining surface accessible residues are in the CDRs and in the interdomain C-terminal region. Figures 3A and 3B show a sequence alignment of the twelve crystal structures, the average relative accessibility, and the 30% accessibility cutoff. Figure 3A shows the alignments relative accessibility for the twelve murine antibody light chains and Figure 3B shows the alignments and relative accessibility for the murine antibody heavy chains.

The surface accessible framework positions were mapped onto a database of unique human and mouse F_V sequences (see lists at the end of this Example). The frequency of particular residues in each of these positions is shown in Table 1. Only residue frequencies higher than 5% are listed.

		Light chain	
5	Position	Human	Mouse
5	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
	5	T 61 L 37	T 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
10	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
	46	P 94	P 82 S 9
	47	G 89	G 71 D 18
	51	K 43 R 31	K 70 Q 13 R 8 T 5
	63	G 91	G 98
	66	D 43 S 25 A 9	D 38 A 26 S 26
15	73	S 96	S 90 I 5
	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
,	87	E 71 D 11 G 7	E 91 D 6
P	111	K 74 R 12 N 6	K 93
00	115	K 54 L 40	K 87 L 5
20	116	R 60 G 33 S 5	R 89 G 9
	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
		Heavy chair	
	Position	Human	Mouse
	118	E 47 Q 46	E 59 Q 29 D 10
25	120	Q 83 T 7	Q 68 K 26
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5
	126	G 54 A 23 P 18	G 36 P 30 A 29
	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
	128	L 61 V 31 F 7	L 96
00	130	K 46 Q 41 E 5	K 52 Q 27 R 17
30	131	P 95	P 91 A 5
	132	G 74 S 16 T 7	G 82 S 17
	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
	143	G 96	G 98
	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
35	160	P 84 S 10	P 89 H 7
	161	G 93	G 71 E 24
•	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
	184	S 70 K 9 P 8	K 42 S 37 T 6
_	186	K 53 Q 22 R 7 N 5	K 83 Q 7
40	187	G 66 S 21 T 5	G 62 S 18 D 10
	1	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
	195	S 91	S 76 A 16
	196	K 65 I 8 T 8 R 5	S 46 K 34 Q 11
	197		T 55 R 26 K 8
45	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11
45	209	A 50 P 21 S 13 T 8	L .
	210	E 46 A 18 D 13 S 9 Z 8 V	T 53 S 43
	212	T 91	·
	222	G 17 D 11 P 10 Y 9 V N 8	D OL V TO

Table 1: Distribution of accessible residues in human VH and VL chain sequences. All of the positions appear to be conserved, which leads to the hyphothesis that immunogeneoity arises from a specific combination of these surface residues. The sequence numbering is explaned in Figures 3A and 3B.

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None of the entire combinations of surface residues in the human sequences are found in the murine sequences and *vice versa* (see lists at the end of this Example). However, the residues in individual positions appear to be conserved (see Table 1). There are few residues which differ significantly between the species;

these are at positions 54 and 91 of the L chain and 168 and 216 of the H chain. Of these positions only position 216 is a non conservative (V to Y) mutation. Differences between human and murine antigenicities are therefore believed to arise from the combinations of residues in these positions.

In order to determine whether the mouse sequences are more distantly related to human F_v sequences than to other mouse F_v sequences, the homology was calculated using a Dayhoff mutation matrix (Dayhoff, M., Barker, W. and Hunt, L. (1983), Meth. Enz. **91**, pp. 524-545). The homology was calculated between all the sequences in a pool of both human and mouse sequence patches made up of the surface accessible residues. The data was then represented as a density map (not shown) in which the sequences are plotted against each other. The density map can be used to discriminate "murine surfaces" from "human surfaces".

B) Reshaping of Antibody N901

In order to test the resurfacing approach suggested by the above analysis, three humanization experiments were set up. 1) Traditional loop grafting (Verhoeyen, M.E., Saunders, J.A., Broderick, E.L., Eida, S.J. and Badley, R.A. (1991), Disease markers 9, pp. 3-4) onto a human F_V framework of known structure (KOL). 2) Resurfacing approach using most similar chain. 3) Resurfacing approach using human sequences with most similar surface residues.

The antibody used was the murine anti-N901 antibody (Griffin et al. (1983), J. Imm. **130**, pp. 2947-2951). The anti-N901 antibody (also referred to herein as the "N901 antibody") is available commercially from Coulter Corporation under the name NKH-1.

The alignment of the light chain sequences and heavy chain sequences in Figures 4A and 4B, respectively, show the original N901 antibody and the sequences used in each of the three approaches outlined here.

Figures 4A and 4B show alignments of sequences generated using the three methods of humanization. Sequences are: 1) Original rodent N901. 2+3) KOL (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391) and reshaped N901 using KOL surface. 4+5) Most homologous sequences, L(KV2F) (Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res., pp. 6499-6513) and H(G36005) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87) and reshaped N901 using these sequences. 6+7) Most homologous with respect to surface residues, L(KV4B) (Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. 3, pp. 6515-6529) and H(PLO123) (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. 168, pp. 229-245), and reshaped N901 using these sequences. The numbering is the same as used in the antibody modelling program ABM (ABM is a trademark for commercial software, Oxford Molecular Ltd., Oxford, U.K.), which is based on structural conservation and not sequence homology as used by Padlan et al. (Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition). The sequence changes which have to be introduced in order to reshape N901 with a given sequence are marked with bars, and back-mutations as determined from F_V models are marked with stars. The sequence homology of a given sequence to N901 is shown in brackets after each sequence.

(1) Classical Humanization

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In classical humanization the rationale is to graft the rodent CDR's onto a framework of known structure, such that CDR-framework interactions can be accurately monitored by homology modelling. The model of the humanized antibody is compared to that of the original rodent antibody, and possible CDR interacting framework residues are back mutated (marked with '*' in alignment) in order to retain the three-dimensional shape of the CDR's. In this example the antibody KOL was used, giving a low homology score of only 77 and 46 in the heavy and light chains respectively.

(2) Most Similar Chain Resurfacing

A database of nonredundant human antibody sequences was compiled from available protein and nucleotide sequences. A total of 164 H and 129 L chains were sampled.

Each of the rodent chains, L and H, were then matched and the most similar human sequence found independently (G36005/KV2F) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87); Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res., pp. 6499-6513). Surface residues, as outlined in Table 1, were then changed in the rodent sequences to match those of the human sequences. Subsequently a model was built of the resurfaced antibody and compared to the model of the original rodent antibody and back mutation of any CDR interacting residues was performed.

(3) Most Similar Surface Replacement According to the Present Invention

This method is identical to the above method, except that the similarity is calculated only over the surface residues outlined in Table 1 above.

The same procedure of surface mutation and subsequent back mutation was performed as in the previous methods. In this case the chosen sequences were PLO123/KV4B (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. **168**, pp. 229-245); Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. **3**, pp. 6515-6529).

The following lists show the surface residue patterns in mouse and human light and heavy chain antibody variable regions. The sequences are ordered on similarity to one another. There are no pattern matches between mouse and human sequences although there are matches within a species.

MOUSE LIGHT CHAIN SURFACE PATCHES

```
:KTSLRPGKGSSDYEKK* (SEQ ID NO: 40)
:KTSLRPGKGSSEYEKK* (SEQ ID NO: 41)
:QTSLRPDKGSSDDEKK* (SEQ ID NO: 42)
:QTSLRPDKGSSDDEKK* (SEQ ID NO: 43)
:QSSLRPDKGSSDPEKK* (SEQ ID NO: 44)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 44)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 45)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 46)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 46)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 47)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 50)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 50)
:QTSLRPDKGSSDPEKK* (SEQ ID NO: 50)
:QTSLRPGKGSSDPEKK* (SEQ ID NO: 51)
:QTSLRPGKGSSDPEKK* (SEQ ID NO: 51)
:QTSLRPGKGSSDPEKK* (SEQ ID NO: 52)
:QTSLRPGKGSSDSEKK* (SEQ ID NO: 53)
:QTSLRPGKGSSDSEKK* (SEQ ID NO: 54)
:PTALRPGKGASDEDKK* (SEQ ID NO: 55)
:VTALRPGKGASDEDKK* (SEQ ID NO: 56)
:VTALRPGKGASDEDKK* (SEQ ID NO: 56)
:VTALRPGKGASDEDKK* (SEQ ID NO: 58)
:QTSLRPGKGSSDPEKK* (SEQ ID NO: 59)
:QTSLRPGKGSSDPEKK* (SEQ ID NO: 60)
:QNSLTPGKGSSDPEKK* (SEQ ID NO: 61)
:VTXVRPGKGDSDBEKK* (SEQ ID NO: 62)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 63)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 66)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 69)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 69)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 69)
:VTKVRPGKGDSDBEKK* (SEQ ID NO: 70)
:VTSVKPGKGDSDBEKK* (SEQ ID NO: 70)
:VTSVKPGKGDSDBEKK* (SEQ ID NO: 71)
                                     1 KV5E$MOUSE
                                     2 PL0101
                                     3 NS1F19L
                                     4 KV5U$MOUSE
                                    5 MUSIGLDD
10
                                    6 PL0220
                                   7 KV5J$MOUSE
                                   8 MUSIGKABB
                              9 MUSIGKABB
9 MUSIGKCLG
10 MUSIGKCLD
11 MUSIGKCLP
12 MUSIGKACM
14 MUSIGKABB
                               15 KVSP$MOUSE
                              16 MUSIGKCHK
17 KV3D$HOUSE
20
                              18 MUSICKAAW
                              19 KV3G$MOUSE
                              20 KV3ESMOUSE
                              21 MUSIGKAAZ
25
                              22 MUSIGKONE
                              23 MUSIGKBA
                              24 KVSASHOUSE
                              25 MUSIGKV
                              26 MUSIGKCNM
                             27 HUSIGKCL1
                             28 KV5B$MOUSE
                             29 MUSIGKCSA .
                             30 MUSIGKCSR
                             31 MUSIGKCST
                                                                                                                                                                                                                      (SEQ ID NO: 69)
(SEQ ID NO: 70)
(SEQ ID NO: 71)
(SEQ ID NO: 72)
(SEQ ID NO: 74)
(SEQ ID NO: 75)
(SEQ ID NO: 76)
(SEQ ID NO: 76)
(SEQ ID NO: 77)
                            32 MUSIGKAB
33 PL0014
                                                                                                       :VTSVKPGKGDSDAEKK*
:VSSVKPGKGDSDAEKK*
:VTSAKPGKGDSDAEKK*
:VSSAKPGKGDSDAEKK*
:VSSAKPGKGDSDAEKK*
:VTSARPGKGDSDAEKK*
:VTSARPGKGDSDAEKK*
:VTSARPGKGDSDAEKK*
:VTSARPGKGDSDAEKK*
:VTFARPGKGDSDAEKK*
:VTTLIPPGKGDSDAEKK*
:VTLIPPGKGDSDAEKK*
:VTLLQPGKGDSDAEKK*
35
                             34 MUSIGRACU
                            35 PS0023
36 N$2MCPL
                             37 MUSIGKADY
                            38 MUSIGECPP
                             39 MUSIGLDS
                            40 MUSIGECHE
                            41 B27887
                            42 H28840
                                                                                                                                                                                                                        (SEQ ID NO: 81)
(SEQ ID NO: 82)
(SEQ ID NO: 83)
(SEQ ID NO: 85)
(SEQ ID NO: 86)
(SEQ ID NO: 87)
(SEQ ID NO: 88)
                            43 KV2G$MOUSE
                            44 C27887
45
                                                                                                                     :Ltllqpgngdsdaekk*
:Vtllqpgkgdsdaeki*
:Vtlpqpggdsdpekk*
                            45 JL0029
                           46 MUSIGKAEH
                            47 PS0074
                                                                                                                  :VTLPOPGKGDSDAEKK*
:VTLPOPGKGDWDAEKK*
                            48 MUSIGKCNY
                            49 MUSIGKCNX
                           50 KV2D$MOUSE
50
                                                                                                                        :VTFLSPGQGDSDAEKK*
                                                                                                                                                                                                                          (SEQ ID NO: 89)
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```
51 MUSIGKADW
                                 : ESSARPGKGDSDAEKK*
                                                         (SEQ ID NO: 90)
      52 KV2A$MOUSE
                                :VTLSSPGQGDSDAEKK*
                                                         (SEQ ID NO: 91)
      53 KVIASMOUSE
                                :VTTAKPEKGDSDVEKK*
                                                         (SEQ ID NO: 92)
      54 F30534
                                : VTTPKPDKGDSDVEKK*
                                                         (SEQ ID NO: 93)
      55 MUSIGKCLO
                                :VTAPRPGKGASSAEKK*
                                                         (SEQ ID NO: 94)
(SEQ ID NO: 95)
      56 G27887
                                :VTAPKPGKGTSSAEKK*
      57 MUSIGVKV3
                                :VTTPKPGKGASSAEKK*
                                                         (SEQ ID NO: 96)
      58 MUSIGKCNA
                                :VSAPKPGKGASSAEKK*
                                                         (SEQ ID NO: 97)
10
      59 S03410
                                :VTAPRSGKGASSAEKK*
                                                         (SEQ ID NO: 98)
      60 B32456
                                :VTAPKSGKGASSAEKK*
                                                         (SEQ ID NO: 99)
      61 PL0013
                                :VTAPKPDKGVSSAEKK*
                                                         (SEQ ID NO: 100)
      62 MUSIGLAET
                                :VTAPKSEKGVSSAEKK*
                                                         (SEQ ID NO: 101)
      63 MUSIGVKV1
                                :FTAPKPGKGASSAEKK*
                                                         (SEQ ID NO: 102)
      64 KV6KSMOUSE
                                 :LTAPKPGRGVSSAEKK*
                                                         (SEQ ID NO: 103)
                                :VTAPKSGKGASSAEKR*
      65 G30560
                                                         (SEQ ID NO: 104)
      66 MUSIGKBO
                                :VSAPKPGKEGSSAEKK*
                                                        (SEQ ID NO: 105)
      67 MUSIGKCNB
                                :VTAPKPRKGASSAEKK*
                                                         (SEQ ID NO: 106)
                                :VTFLSPGQGNSDAELP*
      68 H33730
                                                         (SEQ ID NO: 107)
      69 MUSIGKCPC
                                :VTFLSPGQGNSDEDLP+
                                                         (SEQ ID NO: 108)
                                :VTLSSPQRGDSDAEKK+
      70 KV2CSMOUSE
                                                         (SEQ ID NO: 109)
                                                         (SEQ ID NO: 109)
(SEQ ID NO: 110)
(SEQ ID NO: 111)
(SEQ ID NO: 112)
(SEQ ID NO: 113)
(SEQ ID NO: 114)
(SEQ ID NO: 115)
      71 MUSIGLAV
                                :VTAPKSSKGGSSAEKK*
      72 MUSIGKCNH
                                : QTSPTPGKGSSDPEKK*
      73 KV5R$MOUSE
                                :QISLIPGKGSYDDEKK*
                                 :VTALKSGRGASSAEKK+
      74 KV6ESMOUSE
                                 :VTALKSDKGASSGEKK+
      75 MUSIGKCNI
                                 :VTPPSPGQGDSAAEKK*
      76 MUSIGLDA
                                                         (SEQ ID NO: 116)
(SEQ ID NO: 117)
                                 : VTPPSPGQGDSAREKK*
      77 C26317
      78 PS0073
                                 : VTVRKPGKGDSSDEKK+
                                 : OTSVRLGQGSSDPEKK*
                                                         (SEQ ID NO: 118)
      79 A23986
      80 MUSIGKABW
                                 :KTSLRPWKGSSDSDKK*
                                                         (SEQ ID NO: 119)
                                 : OTDVTQGQGSSQPEXX*
                                                         (SEQ ID NO: 120)
      81 KV5DSMOUSE
                                 :QTAVSQGQGSSQSEKK+
                                                         (SEQ ID NO: 121)
      82 MUSIGE6L
                                 :LTAPRINRGSSDSEKK*
      83 MUSIGECOE
                                                         (SEQ ID NO: 122)
                                 : VTAPSSHRGSSDTEKK*
                                                         (SEQ ID NO: 123)
      84 MUSIGKCKW
                                 :LLSLSPLKGDSDPEKV*
                                                         (SEQ ID NO: 124)
      85 MUSIGLVD
35
                                                         (SEQ ID NO: 125)
                                 : VTAPTPDTGAIKTEKL*
      86 306822
                                 :VTIPTPDTGAIKTEKL*
                                                         (SEQ ID NO: 126)
      87 306821
                                                         (SEQ ID NO: 127)
                                 : AVSPTPDTGAIXTEKL*
      88 MUSIGLAS
                               :AVSPTPDTGAIKTEKL*
                                                         (SEQ ID NO: 128)
      89 MUSIGLAR -
                               : Avsptpdtgviktekl*
                                                          (SEQ ID NO: 129)
      90 LV2B$MOUSE
                                 :AVSPTPDTGAIKTEPS*
                                                         (SEQ ID NO: 130)
      91 MUSIGLAN
```

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HUMAN LIGHT CHAIN SURFACE PATCHES

```
(SEQ ID NO: 131)
(SEQ ID NO: 132)
(SEQ ID NO: 133)
(SEQ ID NO: 133)
(SEQ ID NO: 134)
(SEQ ID NO: 135)
(SEQ ID NO: 136)
(SEQ ID NO: 137)
(SEQ ID NO: 138)
(SEQ ID NO: 138)
(SEQ ID NO: 140)
(SEQ ID NO: 141)
(SEQ ID NO: 142)
(SEQ ID NO: 143)
(SEQ ID NO: 144)
(SEQ ID NO: 144)
(SEQ ID NO: 145)
(SEQ ID NO: 146)
(SEQ ID NO: 147)
(SEQ ID NO: 148)
(SEQ ID NO: 148)
(SEQ ID NO: 149)
                                                                                                      (SEQ ID NO: 131)
(SEQ ID NO: 132)
(SEQ ID NO: 133)
(SEQ ID NO: 134)
(SEQ ID NO: 135)
                 1 LV4ASHUMAN
                                                             :YLPPTPGVIRSTAMKL*
                 2 LV4B$HUMAN
                                                            :YLPPTPGVIRSTAMRL+
                 3 LV4E$HUMAN
                                                            :YLPPTPGLIRSTSMKL*
                 4 LV4DSHUMAN
                                                            :YLPPTPGLIRSTSVKL*
                 5 LV4C$HUMAN
10
                                                            :YLPPTPGVIRSTAEKL*
                                                      :YLPPTPGVIRSTAEKL*
:YLPPTPGVIRSTAGKL*
:YLPATPGVVRSSAGML*
:SLPPSPGKVRSTAEKL*
:SLPPSPGKVRSTAMKL*
:SLPPRPGKVRSSEKL*
:SLPPRPGKVRSSEKL*
:SLPPRPGKVRSSEKL*
:SLPPRPGRVRSSEEL*
:SLPPRPGKVRSSEEL*
:SLPPRPGKVRSSEEL*
:SLPPRPGKVRSSEEL*
:SLPPRPGKVRSSEETL*
                 6 LV5A$HUMAN
                 7 LV7A$HUMAN
                 8 LV2G$HUMAN
                 9 LV2I$HUMAN
               10 N$2RHE
               11 HUMIGLAN
               12 LV1ASHUMAN
               13 LV1B$HUMAN
               14 LV1PSHUMAN
               15 LV1C$HUMAN
               16 A29700
                                                          :SLPPKPGRIRSSTGKL*
20
               17 HUMIGLAM4
                                                            :SLPPKPGKIRSSTGQL*
              18 LV1D$HUMAN
                                                             :SLPPEPGKIRSSTGRL*
                                                                                                        (SEQ ID NO: 148)
(SEQ ID NO: 149)
(SEQ ID NO: 150)
(SEQ ID NO: 151)
(SEQ ID NO: 152)
(SEQ ID NO: 153)
(SEQ ID NO: 154)
(SEQ ID NO: 155)
              19 LV2KSHUMAN
                                                            :SLAPSPGKIRSTAEKL*
               20 LV1ISHUMAN
                                                           : Slpprpgkirsstgnv*
               21 LV2ESHUMAN
                                                           :SLRPSPGKVRSTAEKL*
              22 LV2DSHUMAN
                                                           : SLRPSPGKVRSTADKL*
25
              23 LV2C$HUNAN
                                                           :SLRPSPGKVRSTAENL*
                                                                                                  (SEQ ID NO: 153)
(SEQ ID NO: 154)
(SEQ ID NO: 155)
(SEQ ID NO: 156)
(SEQ ID NO: 157)
(SEQ ID NO: 158)
(SEQ ID NO: 159)
(SEQ ID NO: 160)
(SEQ ID NO: 161)
(SEQ ID NO: 162)
(SEQ ID NO: 163)
(SEQ ID NO: 165)
(SEQ ID NO: 166)
(SEQ ID NO: 166)
(SEQ ID NO: 167)
(SEQ ID NO: 168)
(SEQ ID NO: 169)
(SEQ ID NO: 171)
(SEQ ID NO: 171)
               24 LV2J$HUMAN
                                                          :SLRPSPGKVRSAVEKL*
                                                         :SLPPRPGK-RSSAEKL*
:SLAPSPGKVRSTVERL*
               25 LV1ESHUMAN
               26 LV2B$HUMAN
                                                          :SLAPSPOKIRSTPOKL*
               27 NSINCWN
30
                                                         : Slalspgkvrstaekl*
: Slplsagkvrstaekl*
: Slapspgkvrstaeyl*
: Slpltpglirstaekl*
              28 LV2H$HUKAN
              29 N$3MCG2 .
              30 LV2A$HUMAN
              31 S02083
                                                          :SLPLTPRVIRSTABEL+
              32 HUNIGLAM2
              33 LV6C$HUMAN
                                                          : Flhptpgtdssstekl*
35
              34 LV6D$HUMAN
                                                          : Fllptpgtdsssterl*
              35 LV6ESHUMAN
                                                          : Flhptrvtdssstekl*
              36 LV6B$HUKAN
                                                          : LLPPTPGTMSSSMDKL*
               37 HUMIGLESG
                                                          : VLPLSPHRIRSESENL+
               38 HUNCIGLYC
                                                          : Slapspakfrstaerd*
               39 HUNCIGVLLS
                                                             :VTAPRPGRIRSDPEKK*
40
                                                                                                        (SEQ ID NO: 170)
(SEQ ID NO: 171)
(SEQ ID NO: 172)
               40 HUNIGKAX
                                                             :VTAPRPGRVRSDPEKK*
              41 E30609
                                                             :VTGPRPGRIRSDPEKK*
              42 KV3B$HUMAN
                                                             :VTGPRPGRIRSDPDKK*
                                                       : VTGPRPGRVRSDPEKK*
: VTGPRPGRIRSDPXKK*
                                                                                                    (SEQ ID NO: 173)
(SEQ ID NO: 174)
(SEQ ID NO: 175)
(SEQ ID NO: 176)
              43 G30607
              44 KV3MSHUMAM
45
                                                         :VTAPRPGRIRSESERK*
:VTGPSRGRIRSDPEKK*
              45 KV3H$HUMAN
                                                          :VTGPSRGRIRSDPEKA-
:VTVPRPSRIRSESERK* (SEQ ID NO: 177)
:VTAPGPGRIRSESERK* (SEQ ID NO: 178)
:QTSVRPGRVRSDPERK* (SEQ ID NO: 179)
.OTSVRPGRVRSDPERK* (SEQ ID NO: 180)
              46 KV3K$HUMAN
           47 KV3FSHUMAM
              48 B26555
               49 KV1Q$HUMAN
             50 KV1WSHUHAN
```

```
51 KV1MSHUMAN
                                                         (SEQ ID NO: 181)
                                :QTSVRPGKVRSDPEKK*
     52 KV1R$HUMAN
                                                         (SEQ ID NO: 182)
                                :QTSVRPGKVRSEPEKK*
                                                         (SEQ ID NO: 183)
(SEQ ID NO: 184)
(SEQ ID NO: 185)
(SEQ ID NO: 186)
     53 KV1F$HUMAN
                                :QTSVRPGKVRSEPDKK*
     54 KV1G$HUMAN
                                :QTSVRPGKVRAEPEKK*
     55 KV1K$HUMAN
                                :QTSVRPGKVRSBPZKK*
     56 KV1DSHUMAN
                                :QTSVRPGKVRSDPBKK*
                                                         (SEQ ID NO: 187)
     57 KV1H$HUMAN
                                :QTSVRPGQVRSDPERK*
                                                         (SEQ ID NO: 188)
     58 KV1B$HUMAN
                                : OTSVRPGKVRSHPEKK*
                                                         (SEQ ID NO: 189)
10
     59 B27585
                                :OTSVRPGNVRSDPDKK*
                                                         (SEQ ID NO: 190)
     60 NSIREIA
                                :QTSVRPGKVRSDPEKT*
                                                         (SEQ ID NO: 191)
                                :QTSVRPGTVRSEPEKK*
     61 KV1XSHUMAN
                                                         (SEQ ID NO: 192)
     62 KV1L$HUMAN
                                :QTSVRPEKVRSEPDKK*
                                                         (SEQ ID NO: 193)
     63 IMGL38
                                :QTSVRPGKVRSESDKK*
                                                         (SEQ ID NO: 194)
                                :QTSVRPGEVRSEPDKK*
     64 A27585
15
                                                         (SEQ ID NO: 195)
     65 KV1NSHUMAN
                                :QTSVRPGBVRSBPZRK*
                                : OTSVSPGKVRSDPEKK*
                                                         (SEQ ID NO: 196)
     66 KV1C$HUMAN
     67 KV1VSHUMAN
                                : QTSVRPGKVNSDPEKK*
                                                         (SEQ ID NO: 197)
                                :QTSVRPGKVRSDPDTK*
     68 KV1TSHUMAN
                                                         (SEQ ID NO: 198)
                                : QTSVRPKKVRSDP2KK*
                                                         (SEQ ID NO: 199)
     69 KV1USHUMAN
                                                          (SEQ ID NO: 200)
                                :QTSVRPKKVRFDPEKK*
     70 KV1ASHUMAN
20
                                                          (SEQ ID NO: 201)
                                :QTSVRSGKVRSEPETK*
     71 KV1S$HUMAN
                                                          (SEQ ID NO: 202)
                                : VTNLRPGKVRSDAEKK*
      72 KV4A$HUMAN
                                                          (SEQ ID NO: 203)
                                :VTDLRPGKVRSDAEKK*
      73 KV4C$HUMAN
                                                          (SEQ ID NO: 204)
                                :QTSVSPGNIRSESDKK*
      74 HUMIGK2A1
                                                          (SEQ ID NO: 205)
                                :KTSVTPGKFRSEPEKK*
      75 HUMIGKBA
25
                                                          (SEQ ID NO: 206)
                                :VTLLPPGRVRSDAEKK*
      76 HUMIGKBC
                                                          (SEQ ID NO: 207)
                                :VTLLPPGEVRSDAEKK*
      77 KV2BSHUMAN
                                                          (SEQ ID NO: 208)
                                : VTLPPPGZVRSDAERK*
      78 KV2D$HUMAN
                                                          (SEQ ID NO: 209)
                                :VTLPPPGZVRSBAZNK*
      79 KV2C$HUMAN
                                                          (SEQ ID NO: 210)
                                :VTLPPPQQVRSDAEKK+
      80 KV2ESHUMAN
                                :VTLPPPGQVTSDAEKK*
                                                          (SEQ ID NO: 211)
      81 S03876
30
                                :VTLPPAGQVRSDAEKR*
                                                          (SEQ ID NO: 212)
      82 KV2A$HUMAN
                                                          (SEQ ID NO: 213)
                                :ALSPSSGQSSSASERL*
      83 HUMIGLAMS
```

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HOUSE HEAVY CHAIN SURFACE PATCHES

```
1 MUSIGHIT
                                : EKVGGLQPGRGTPGKASRGDSQRPES*
                                                                  (SEQ ID NO: 214)
       2 MUSIGHIU.
                               : EKVGGLQPGRGTPGKVSRGDSQRPES*
                                                                  (SEQ ID NO: 215)
10
       3 MUSIGHIV
                                : EKVGGLQPGTGAPGKASRGDSQRPES*
                                                                  (SEQ ID NO: 216)
       4 MUSIGHYM
                               : EKVGGLQPGRGTPGKASKGNSQRAES*
                                                                  (SEQ ID NO: 217)
       5 PU0003
                               : EKMGGLQPGRGTPGKASKGNSQRAES*
                                                                  (SEQ ID NO: 218)
      6 MUSIGHFO
                               : EKVGGLQPGRGTPGKASKGTSQRAES*
                                                                  (SEQ ID NO: 219)
         A30515
                               : EKVGGLQPGRGTPGKASKGTSQRAET.
                                                                  (SEQ ID NO: 220)
      8 PL0018
                               : EKVGGLKPGRGTPGKASKGTSQRAET*
                                                                  (SEQ ID NO: 221)
      9 MUSIGHFK
                               : ENVGGLQPGRGTPGKASKGTSQRAET*
                                                                  (SEQ ID NO: 222)
     10 MUSIGHPQ
                               : EKVGGLQSGRGTPGKASKGTSQRAET*
                                                                 (SEQ ID NO: 223)
(SEQ ID NO: 224)
     11 PU0001
                               : EKVGGLQSGRGTPGKASKGTSQRAES*
     12 E30540
                               : EKVGGLQPGRGTPGKASKGISQRAER*
                                                                  (SEQ ID NO: 225)
     13 HV17$MOUSE
                               : EKVGGLQPGRGTPGKSAKGBSZRAQS*
                                                                  (SEQ ID NO: 226)
     14 MUSIGHLN
                               : EKVGGLQPGSGTPGKASKGNSQRAES+
                                                                  (SEQ ID NO: 227)
     15 MUSIGHKG
                               : EKVGGLQPGSGTPGKASKGSSQRAES+
20
                                                                  (SEQ ID NO: 228)
     16 PU0004
                               : EKVGGLQPGRGTPRKASKGNSQRAES+
                                                                  (SEQ ID NO: 229)
     17 MUSIGHKJ
                               : EKMGNLQPGSGTPGKASKGNSQRPDS+
                                                                  (SEQ ID NO: 230)
     18 HV56SMOUSE
                               : EKVGGLKPGKGTPEKDSKGNARRSET+
                                                                  (SEQ ID NO: 231)
     19 C27888
                               : EKVGGLKPGKGAPEKDSKGNARRSET*
                                                                  (SEQ ID NO: 232)
     20 MUSIGHAAP
                               : EKVGGLKPGKGTPERDSKGNARRSET+
                                                                 (SEQ ID NO: 233)
     21 PH0097
                               : DKVGGLKPGKGTPEKDSKGNAKRSET+
                                                                 (SEQ ID NO: 234)
25
     22 E27888
                               : DKVGGLKPGKGTPEKDSKGNAKKSET*
                                                                 (SEQ ID .NO: 235)
     23 MUSIGHJB
                               : DKVGGLKPGKGTPDKDNKGNAKKSET+
                                                                 (SEQ ID NO: 236)
     24 MUSIGHADL
                               : EKVGGLTPGKGTPEKDSKGNGRRSET+
                                                                 (SEQ ID NO: 237)
     25 A27888
                               : ENVGGLKPGKGTPEKDSKGNDRRSET*
                                                                 (SEQ ID NO: 238)
     26 H27887
                               : ENVGGLKPGKGTPEKDSKGNDKRSET+
                                                                 (SEQ ID NO: 239)
     27 B27888
                               : Emvgglkpgkgtpekdskgnakrset•
                                                                 (SEQ ID NO: 240)
     28 B27889
                               : BOVGGLKPGKGTPEKDSKGHAKKSET*
                                                                 (SEQ ID NO: 241)
     29 D27889
                               : EQVGGLKPGKGTPEKDTKGNAKKSET+
                                                                 (SEQ ID NO: 242)
     30 HV55$MOUSE
                               : BOVGGLKPGKGAPEKDTKGNAKKSET+
                                                                 (SEQ ID NO: 243)
     31 MUSIGHAGT
                               : EKVGGLOPGKGTPEKDSKGNAKKSET+
                                                                 (SEQ ID NO: 244)
     32 MUSIGVH50
                               : EKVGGLQPGKGTPEKDTKGNAKKSET*
                                                                 (SEQ ID NO: 245)
     33 MUSIGHIW
                               : EKVGGLOPGRGTPEKDTKGNAKKSET+
                                                                 (SEQ ID NO: 246)
                               : EKVGGLOPGKGSPEKDSKGNAKKSET*
     34 MUSIGHAGE
35
                                                                 (SEQ ID NO: 247)
     35 PH0098
                               : DKMCGLKPGKGTPEKDSKGNAKOSET+
                                                                 (SEQ ID NO: 248)
     36 MUSIGHIM
                               : EQVGGLQPGKGTPDKDSKGNAKKSET+
                                                                 (SEQ ID NO: 249)
     37 MUSIGHAGY
                               : EKVGGLQPGKGTPEKDSKGNAEKSET+
                                                                 (SEQ ID NO: 250)
     38 MUSIGHDET
                               : EQVGDLKPGKGTPEKDTKGNARRSET*
                                                                 (SEQ ID NO: 251)
     39 D27888
                               : ENVGDLKPGKGAPEKDSKGNARRSET+
                                                                 (SEQ ID NO: 252)
     40 MUSIGHIP
                               : EQVGGLQPGKGTSDKDSKGNAKKSET+
                                                                 (SEQ ID NO: 253)
40
     41 MUSIGHAGS
                               : EQVGGLQPGKGTPEKDSKGNAKKSGT+
                                                                 (SEQ ID NO: 254)
     42 HV16$MOUSE
                               : DQVGGLQPGKGTPEKDTKGNPKRSET+
                                                                 (SEQ ID NO: 255)
     43 B34871
                               : DQVGGLQPGQGTPEKNTKGNPKRSDT+
                                                                 (SEQ ID NO: 256)
                               : EKVGGLOPGKGTSEKDIKGKAKKSET*
                                                                 (SEQ ID NO: 257)
     44 PH0094
     45 PH0096
                               : DKVGGLKPGKRTPEKDNKGNAKKSET+
                                                                 (SEQ ID NO: 258)
     46
       MUSIGVH62
                               : DKVGGLKLGKGTPEKDTKGNAKKSET+
                                                                 (SEQ ID NO: 259)
45
     47 MUSIGHAGR
                               : EKVGGLQPGKGTPEKDSKGNAHTSET*
                                                                 (SEQ ID NO: 260)
     48 HV58$MOUSE
                               : EHVGGLAPGKGTPEKDSKGNAGRSET*
                                                                 (SEQ ID NO: 261)
     49 H27888
                               : EQVGGLQPGHGTPEKDTTGHAKRSET+
                                                                 (SEQ ID NO: 262)
     50 HV34$MOUSE
                               : EKEGGLOPGKGTPEKESKGDSKRAET*
                                                                 (SEQ ID NO: 263)
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51 HV33$MOUSE
                                : EKEGGLQPGKGTPEKESKGDSKRPET+
                                                                  (SEQ ID NO: 264)
      52 MUSIGHZAB
                                : EKEGGLQPGKGSPEKESKGDSKRAET*
                                                                   (SEQ ID NO: 265)
      53 NS4FABH
                                : EKDGGLQPGKGTPEKDSKGDSKRVEM*
                                                                  (SEQ ID NO: 266)
      54 I27888
                                : EQVGGLKPGRGTPEKDTTGDAQRSET*
                                                                   (SEQ ID NO: 267)
      55 G27888
                                : EQVGGLKPGRGTPEKDTTGNAKGSET*
                                                                  (SEQ ID NO: 268)
10
      56 HV59$MOUSE
                                : EKVGGSKPGKGTPEKDSKGNAKTSET*
                                                                  (SEQ ID NO: 269)
      57 MUSIGHOE
                                :SDQGGLKPGKGTPEKDTKGNARRSES*
                                                                   (SEQ ID NO: 270)
      58 N$2FVWH
                                : EKIGGLOPGKGDPGKPSKDNAKRSET*
                                                                   (SEQ ID NO: 271)
                                : EKLGGLOPGKGDPGKPSKDNAKRSET*
      59 MUSIGHJT
                                                                  (SEQ ID NO: 272)
      60 MUSIGHLY
                                : EKLGGLQPGKGDPGKPFKDNAKRSET*
                                                                   (SEQ ID NO: 273)
      61 S06816
                                : EKLGGLQPGKGDPGKLMKENAKRSET+
                                                                  (SEQ ID NO: 274)
      62 506817
                                : ENLGGLQPGKGDPGKLKXENAKRPET*
                                                                  (SEQ ID NO: 275)
      63 MUSIGHAAI
                                : EKLGGLQPGNGDLGKPSKDNAKRSET+
                                                                   (SEQ ID NO: 276)
      64 HV42$MOUSE
                                : EKLGPLQLGKGDPGKPSKDDAKRSET*
                                                                   (SEQ ID NO: 277)
                                                                   (SEQ ID NO: 278)
      65 MUSIGHAAL
                                : EQLGGLQPGGGTPGKPSKDNDKRSET*
      66 MUSIGHABO
                                : EQLGGLQPGGGTPGKASKDNDKRSET*
                                                                   (SEQ ID NO: 279)
      67 MUSIGHEG
                                : EQVGGLKARKGTPEKDTTGNAKRSET*
                                                                   (SEQ ID NO: 280)
      68 MUSIGHWN
                                : EMVGVLEPGKGTPEKRQEGNAKRSET*
                                                                   (SEQ ID NO: 281)
20
      69 MUSICKCLT
                                : EQVGGLQPKKGSPGKDSKDDSQKTET*
                                                                   (SEQ ID NO: 282)
      70 MUSIGHZAE
                                : EQVGGLQPKKGSPGKDSKDDSQKTER*
                                                                   (SEQ ID NO: 283)
      71 MUSIGHAAD
                                : QQVPELKPGRGTPGKEDKGTSARNDT*
                                                                   (SEQ ID NO: 284)
      72 MUSIGHAAW
                                :QQVPELKPGKGTPGKDDKGTSAKNET*
                                                                   (SEQ ID NO: 285)
      73 MUSIGHAMA
                                :QQVPELKPGKGTPGKDDKGTSAKNEM*
                                                                   (SEQ ID NO: 286)
                                                                   (SEQ ID NO: 287)
      74 MUSIGHXZ
                                : QQKPELKPGKGSPGQEKKGTSSTSET*
25
      75 A30502
                                : EQQPELXPGKGTPGQEKKGKSSTSES*
                                                                   (SEQ ID NO: 288)
      76 MUSIGHAAG
                                : EQQPELRPGKGTPGQEKKGKSSTSES*
                                                                   (SEQ ID NO: 289)
                                : EQQPELKPGKGTPGQEKKGKSSASES*
                                                                   (SEQ ID NO: 290)
      77 B30502
      78 MUSIGHADG
                                : EQQPELKPGKGTPGKQKKGKSSTSES*
                                                                   (SEQ ID NO: 291)
                                                                   (SEQ ID NO: 292)
      79 MUSIGHPV
                                : ECOPELKPGKGTHGKOKKGKSSTSES*
                                                                   (SEQ ID NO: 293)
      80 MUSIGHAANA
                                : EQQPELKPGKGSHGKQKKGKSSTSES*
      81 MUSIGHZR
                                : EQQPELKPGKGSHGKQKKGKSSASES*
                                                                   (SEQ ID NO: 294)
30
                                                                   (SEQ ID NO: 295)
                                : eqqpelkpgkgthgkqkkgksstfes*
      82 MUSIGHAI
      83 MUSIGHALA
                                : EQQPELKPGKGTHGKQKQGKSSTFES*
                                                                   (SEQ ID NO: 296)
      84 PL0011
                                : EQOPELKPGKGTHGKEKKDKSSTSES*
                                                                   (SEQ ID NO: 297)
                                                                   (SEQ ID NO: 298)
      85 MUSIGKCLS
                                : BOOAELKPGKGSHGKOKKGKSSTSES*
                                : EQQPELEPGEGTHGKQEKSNSSTSES*
                                                                   (SEQ ID NO: 299)
      86 MUSIGHADY
                                                                   (SEQ ID NO: 300)
(SEQ ID NO: 301)
                                : OOOAELRPGKGAPGQEKKGKSSTSES*
      87 MUSIGHWVX
35
      88 MUSIGRADO
                                : QQQAELRPGKGAPGQEKKGKSSTSD8*
                                : QQQAELRPGKGVPGQEKKGKSSTSDS*
                                                                    (SEQ ID NO: 302)
      89 MUSIGHVBM
                                                                   (SEQ ID NO: 303)
                                : OOOPELKPGKGAPGKGKKGKSSTSES*
      90 A24672
                                                                   (SEQ ID NO: 304)
                                : QQQPELKPGKGAPGKGKKDKSSTSES*
      91 MUSIGHJG
                                                                   (SEQ ID NO: 305)
                                : ECOPEAKPGKGTHGKQKKGKSSTSDS*
      92 JL0044
                                                                   (SEQ ID NO: 306)
(SEQ ID NO: 307)
                                : QQQAELKPGKGTHGKEKKDKSSTSDS*
      93 MUSIGHBA
40
                                : QQQAELRPGKGAPGQGKKGKSSTSES*
      94 MUSIGRAGP
                                                                    (SEQ ID NO: 308)
                                : QQQAELKPGRGTPGQEKKGKSSTSES*
      95 MUSIGHVBK
                                                                    (SEQ ID NO: 309)
                                : EQQAELRAGKGTPGQEKKGKSSTSES*
         A36194
      96
                                                                    (SEQ ID NO: 310)
                                : EQQAELRPGKGTPGQEKKGTSSTSES+
      97
         MUSIGHVBJ
                                : QQQAELRPGKGTPGHEKKGTSSTSES*
                                                                    (SEQ ID NO: 311)
      98 MUSIGHADV
                                : QQQAELKPGKGTPGHEKKGTSSTSES*
                                                                    (SEQ ID NO: 312)
      99
         MUSIGHAAT
                                : QQQAELRPGKGTPGHENKGTSSTSES*
                                                                    (SEQ ID NO: 313)
     100 MUSICHJL
```

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5
      101 MUSIGHABM
                                  : QQQAEVRPGKGTPGHEKKGTSSTSES*
                                                                     (SEQ ID NO: 314)
      102 MUSIGHFU
                                  : QQQAELKPGKGTPGHENKGTSSTSES+
                                                                     (SEQ ID NO: 315)
      103 MUSIGHZZB
                                  : QQQAELRPGKGTPGQQKKGKSSASES*
                                                                     (SEQ ID NO: 316)
     104 HV06$HOUSE
                                  : HQQAELKPGKGTPGQQKKGKSSTSES*
                                                                     (SEQ ID NO: 317)
(SEQ ID NO: 318)
     105 MUSIGHRD
                                 : EQQVELRAGKGTPGQEKKGKSSTSES*
10
     106 MUSIGHVBH
                                 : EQQAELRPGKGTPGQEKQGTSSTSES*
                                                                     (SEQ ID NO: 319)
     107 HV01SMOUSE
                                 : EQQAELRPGKGTPGHDNKGTSSTSES+
                                                                     (SEQ ID NO: 320)
     108 MUSIGHADN
                                 : QQQAEVRPGKGTPGHEKKGRSSTSES*
                                                                     (SEQ ID NO: 321)
(SEQ ID NO: 322)
     109 HVOSSHOUSE
                                 : QQQAELRPGKGTPGQQKKDKSSTSES*
     110 MUSIGHAEF
                                 : QQQAELKPGKGTPGQQKKDKSSTSES+
                                                                     (SEQ ID NO: 323)
     111 MUSIGHAAN
                                 : QQQAELKPGKGTPGQQKKDKSSTSDS*
                                                                     (SEQ ID NO: 324)
     112 MUSIGHAAB
                                 : QQQAELKPGKGSPGQQKKDKSSTSES*
                                                                     (SEQ ID NO:
                                                                                  325)
     113 C30560
                                 : QHQAELKPGKGTPGQQKKNKSSTSES*
                                                                     (SEQ ID NO: 326)
     114 PS0024 -
                                 : QQQAELKPGKGTPGQQNKDKSSTSES*
                                                                     (SEQ ID NO: 327)
     115 MUSIGHRG
                                 : EQQAELRAGKGIPGQEKKGKSSTSES*
                                                                     (SEQ ID NO: 328)
(SEQ ID NO: 329)
     116 MUSIGHAAR
                                 : QQQAELKPGKGTPGQEKKSKSSTSES+
     117 MUSIGHLX
                                 : QQQSELKPGKGTPGQEKKSKSSTSES*
                                                                     (SEQ ID NO: 330)
20
     118 HV04$MOUSE
                                 : QQQTELKPGKGTPGQEKKSKSSTSES+
                                                                     (SEQ ID NO: 331)
                                 : EQQAELRTGKGTPGQERKGKSSTSES+
     119 MUSIGHVBG
                                                                     (SEQ ID NO: 332)
     120 MUSIGHNOX
                                 : QQQAELKPGKGTPGQQKKDKSSTFES*
                                                                     (SEQ ID NO: 333)
     121 MUSIGHAAR
                                 : EQQAELRPGTGAPGQEKKGKSSTSES*
                                                                     (SEQ ID NO: 334)
     122 HV15$MOUSE
                                 : QQQPEVRPGKGTHAKQKKGKSSTSES*
                                                                     (SEQ ID NO: 335)
(SEQ ID NO: 336)
     123 MUSIGHAAU
                                 : QQQPEVRPGKDTHAKQKKGKSSTSES*
     124 MUSIGHVBO
                                 : QQQAELKPGKGTPEQEKKGKSSTSES*
25
                                                                     (SEQ ID NO: 337)
     125 A26405
                                 : EQQTELRAGKGTPGQEKKGRSSTSZA+
                                                                     (SEQ ID NO: 338)
     126 HV10$MOUSE
                                 : QQQAELKPGKGTPGREKKSKPSTSES*
                                                                     (SEQ ID NO: 339)
     127 MUSIG3B44
                                 : QQQSELKPGKGTPGREKKSKPSTSES*
                                                                     (SEQ ID NO: 340)
     128 MUSIG3B62
                                 : QQRAELKPGKDTPGREKKNKPSTSES*
                                                                     (SEQ ID NO: 341)
     129 HV09$MOUSE
                                 : QQQAELKPGKGTPGREKKSTSSTSES*
                                                                     (SEQ ID NO: 342)
(SEQ ID NO: 343)
     130 MUSIGKCLP
                                 : QQQAELKPGKGTPGQEKKSTSSTSDS*
     131 MUSIGBH
                                 : QQQAELRPGKGTPIQQKKDKSSTSES*
                                                                     (SEQ ID NO: 344)
     132 HV11$MOUSE
                                 : QQQAEPKPGKGTPGREHRSKPSTSES*
                                                                     (SEQ ID NO: 345)
(SEQ ID NO: 346)
     133 MUSIGHMC
                                 :QQQAELRPGKGALGQEKKGKSSTSDS+
     134 MUSIGHAGN
                                 :QQQPEVKPGKGAPGKGNTDKSSTSES*
                                                                     (SEQ ID NO: 347)
     135 MUSIGHRY
                                 : Equaevragegspeqeeeegestses+
                                                                     (SEQ ID NO: 348)
(SEQ ID NO: 349)
     136 MUSIGHVAD
                                 :QQLAELKPGKGTPGHEKKGI99T9E9*
35
     137 MUSIGHVAP
                                 : QQQAELKPGKGKPEQEKKGT99T5E9*
                                                                     (SEQ ID NO: 350)
     138 PL0012
                                 : QQQPELXPGKGRNGKENKGKSSTSES*
                                                                     (SEQ ID NO: 351)
     139 MUSIGGVD2.
                                 : QQQTELRPGRGTTGQERKGK9STSES*
                                                                     (SEQ ID NO: 352)
(SEQ ID NO: 353)
     140 806824
                                 : QHQARLKPGKGTPGHENKVTSSTSES*
     141 MUSIGHTS
                                 : EQQAELRAGKGTPGQEQKAKSSTSES*
                                                                     (SEQ ID NO: 354)
     142 MUSIGHAAM
                                 :QQQAELKPGKGTPGQQKTGTSSTTES*
                                                                     (SEQ ID NO: 355)
     143 MUSIGHHS
                                 : QQQAELKPGKGNPGQEKKSTSSASES*
                                                                     (SEQ ID NO: 356)
     144 MUSIGHAXA
                                 : EQOTVLRPGKGTPGQQKKGTSATNES*
                                                                     (SEQ ID NO: 357)
     145 HV50$MOUSE
                                 : QQLTELKPGMGTPGQEKKSKSSTSES*
                                                                     (SEQ ID NO: 358)
                                 : OOQSVLRPGKGTPGQEKKGTSSTSKS*
     146 MUSIGHVBP
                                                                     (SEQ ID NO: 359)
     147 PH0100
                                 : LQQPVLKPGKGSHGKQKKDKSSTSES*
                                                                     (SEQ ID NO: 360)
     148 MUSIGHAYA
                                 : EQQPETKPGKGTLGKQKKSKSSTSES*
                                                                     (SEQ ID NO: 361)
     149 MUSIGHCP2 -
                                 :QQQAELKPGQGTPGQEKKKSSTPEP*
45
                                                                     (SEQ ID NO: 362)
```

150 MUSIGHDZ

55

: EQQAELRPGKGMPEQPKQGTSSTSET*

(SEQ ID NO: 363)

```
151 MUSIGHNPI
                                 : EQQAELRPGKGNPEQPKQGTSTTSET+
                                                                   (SEQ ID NO: 364)
      152 506823
                                 : EQQAELKPGKGNPEQPKQGTSSTSET*
                                                                   (SEQ ID NO: 365)
      153 MUSIGHASA
                                 : EQQAELKPGKGNPEQPKQDTSSTSET+
                                                                   (SEQ ID NO: 366)
      154 S03484
                                 : EQQAELKPGKGNPEQPKQGTSSTSGT+
                                                                   (SEQ ID NO: 367)
      155 MUSIGHVAA
                                 : EQQAEVKPGKGNPEQPKQGTSSTSET*
10
                                                                   (SEQ ID NO: 368)
      156 MUSIGHNPD
                                 : EQQAELRPGKGNPEQPKQVTSSTSET*
                                                                   (SEQ ID NO: 369)
      157 MUSIGHNPB
                                 : EQQAELRPGKGNPEQPKQITSSTSET*
                                                                   (SEQ ID NO: 370)
      158 MUSIGHEC
                                 : EQQAELRPGRGNPEQPKQVTSSTSET*
                                                                   (SEQ ID NO: 371)
      159 MUSIGHNPC
                                 : EQQAELRPGRGNPEQPKHVTSSTSET+
                                                                   (SEQ ID NO: 372)
(SEQ ID NO: 373)
      160 MUSIGHNPF
                                 : EQQAELRPGKGNTEQPKQVTSSTSET*
      161 MUSIGHNPE
                                 : EQQAELKPGKGNTEQPKLITSSTSET+
                                                                   (SEQ ID NO: 374)
      162'A27635
                                 :TGQAELRPGKGAPEQGKKGKSSTSDR+
                                                                   (SEQ ID NO: 375)
      163 MUSIGHXW
                                 :QYQAELRPGKGTPRQQKKGKSSTSES+
                                                                   (SEQ ID NO: 376)
      164 MUSIGHIZA
                                 : QQQAVLRHGKGTHGQEKKGKSSTSES*
                                                                   (SEQ ID NO: 377)
      165 MUSIGHEH
                                 : QQQTKLGPGRGTPGQGRKGKSSTSGS+
                                                                   (SEQ ID NO: 378)
      166 MUSIGHRH
                                 : Eqqaelragkgtpgqekkgkssvypa+
                                                                   (SEQ ID NO: 379)
                                 : EQQAELKAGKGTPGQQKQGESTRSET+
      167 HV00$MOUSE
                                                                   (SEQ ID NO: 380)
      168 N$1F19H
                                 : QQKAELAASKGTPGQEKKGRSSTSES*
20
                                                                   (SEQ ID NO: 381)
      169 MUSIGHZAD
                                 : QQQTELRPGKGTPGQEKRGKSSNLRL*
                                                                   (SEQ ID NO: 382)
      170 B30515
                                 : EKVGGLQGSSFDPGKASKGTSQRAET+
                                                                   (SEQ ID NO: 383)
      171 MUSIGHER
                                 : EQQADLKLGKGNPEQPKLATPSTSET+
                                                                   (SEQ ID NO: 384)
      172 E27889
                                 : EQVGGLKPGKGTPDKSDVKDNAKSET*
                                                                   (SEQ ID NO: 385)
      173 MUSIGHAAC
                                 : DQQPDLKPSSGSPGHPSKSTSKTTET*
                                                                   (SEQ ID NO: 386)
      174 HV61$MOUSE
                                 : DQQPDLKPSSGSPGNPSKSTSKTTET*
                                                                   (SEQ ID NO: 387)
25
      175 MUSIGVHR2
                                 : DQQPDLKPSSGSPGNPSKSTSKTAET*
                                                                   (SEQ ID NO: 388)
      176 PL0100
                                 : DQQPGLKPSSGSPGNPSKSTSKTTET*
                                                                   (SEQ ID NO: 389)
      177 MUSIGHAAO
                                 : DQQPGLKPSSGSPGNPSKNTSKTTET*
                                                                   (SEQ ID NO: 390)
      178 MUSIGHGA6
                                 : DQQPGLKPSSGSPGDPSKTTSKTTET+
                                                                   (SEQ ID NO: 391)
                                 : DOOPGLEPSSGSPGNPSKTTSKTTET+
     179 MUSIGHJY
                                                                   (SEQ ID NO: 392)
      180 MUSIGHGA1
                                 : DHQPGLKPSSGSPGNPSKNTSKTTET*
                                                                   (SEQ ID NO: 393)
     181 MUSIGHXX
                                 : DQQPGLKPSSGSPGNPSRSTSKTTET+
30
                                                                   (SEQ ID NO: 394)
     182 HV62$MOUSE
                                 : DQQPGLKPSAGSPGNPSKSTSKTAET*
                                                                   (SEQ ID NO: 395)
     183 MUSIGHAAGA
                                 : EQQPGLKPSSGSPGNPSRSTSKTSET*
                                                                   (SEQ ID NO: 396)
     184 MUSIGHGA5
                                 : DOOPGLKPSSGSPGNPSKNTSKTIET*
                                                                   (SEQ ID NO: 397)
     185 MUSIGHGA4
                                 : DOOPGLKPSSGSPGDPSKNTSKTPET*
                                                                   (SEQ ID NO: 398)
     186 MUSIGHAGI
                                 : EQQPSLKPSSGSPGNPSKSTSKTTET+
                                                                   (SEQ ID NO: 399)
     187 PL0102
                                 : DQQPGLKPSSGSPGNPSKNTSETTET*
                                                                   (SEQ ID NO: 400)
35
     188 HV46$MOUSE
                                 : DQQPGLKPSSGSPGNPSKNTSETTZT*
                                                                   (SEQ ID NO: 401)
     189 MUSIGHZT
                                 : EQQPSLKPSSGSPGNPSKSTSKTSET+
                                                                   (SEQ ID NO: 402)
     190 MUSIGHAGD
                                 : ECOPSLKPSSGSPGNPSKSTSRTTET+
                                                                   (SEQ ID NO: 403)
     191 MUSIGHAGO
                                 : EQQPSLKPSSGSPGNPSKSTSKTAET+
                                                                   (SEQ ID NO: 404)
     192 MUSIGAM32
                                 : DQQPDLKPSSGFPGNPSKSTSKTTET*
                                                                   (SEQ ID NO: 405)
     193 MUSIGHAFX
                                 : POOPSLEPSSGSPGKPSKSTSKTNET*
                                                                   (SEQ ID NO: 406)
                                 : EQQPSLKPSSGSPGNPSKSTYKTSET+
40
     194 MUSIGHAGE
                                                                   (SEQ ID NO: 407)
     195 MUSIGHAGE
                                 : ECOPSLIKPSSGSPGNPSKSTSTTSET*
                                                                   (SEQ ID NO: 408)
     196 MUSIGHAGC
                                 : EQQLSLKPSSGSPGNPSKSTSKTTET*
                                                                   (SEQ ID NO: 409)
     197 MUSIGHAAM
                                 :QQQPGLKPSPGPPGKPSQSTSKTTET*
                                                                   (SEQ ID NO: 410)
     198 HV43$MOUSE
                                 : QOKPGLAPSSGSPGKSTKSNSKQTDT*
                                                                   (SEQ ID NO: 411)
     199 MUSIGMUV1
                                 : QOKPGLAPSSGSPGKSAKSNSKQTDT*
                                                                   (SEQ ID NO: 412)
     200 MUSIGHAEI
                                 : QOKPGLAPSGSPGKSAMSNSKQTDT*
                                                                   (SEQ ID NO: 413)
45
                                 : QQKPGLAPSSGSPGXSAISMSKQTDT+
     201 MUSIGHBP
                                                                   (SEQ ID NO: 414)
     202 MUSIGHZZA
                                 : QQKPGLQPSSGSPGKAAISNSKQSNT*
                                                                   (SEQ ID NO: 415)
     203 MUSIGMUV2
                                 :QOKPGLQPSSGSPGKAAISNSKQANT*
                                                                   (SEQ ID NO: 416)
                                 : QQKPVLAPSSGSPGKSAMSNSKQIDT*
     204 A32456
                                                                   (SEQ ID NO: 417)
      205 MUSIGHMB
                                 : QQKPSLQPSSDSPGKAAMSHSKQADT*
                                                                   (SEQ ID NO: 418)
```

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HUMAN HEAVY CHAIN SURFACE PATCHES

```
1 HUMIGHVS
                                : ERVGDLEPGRGIPGKAPKGDSKKIET*
                                                                  (SEQ ID NO: 419)
       2 HUMIGHVR
                                : ERVGDLEPERGIPGKAPKGDSKKIET*
                                                                  (SEQ ID NO: 420)
       3 H36005
                                : EQVGGLKPGRGTPGKAPKGDSKKTET*
10
                                                                  (SEQ ID NO: 421)
       4 PL0122
                                : EQVGGLQPGKGTSGKASKGDSKKTET*
                                                                  (SEQ ID NO: 422)
       5 HV3D$HUMAN
                                : EQLGGLQPGRGTPGKBSKGDSKRAET*
                                                                  (SEQ ID NO: 423)
       6 HUMIGHAT
                                : EQLGGLQPGRGTPGKDSKGNSKRAET+
                                                                  (SEQ ID NO: 424)
         B34964
                                : EQLGGLQPGRGTPGKDSRGNSKRAET*
                                                                  (SEQ ID NO: 425)
         A34964
                                : EQVGGLQPGRGTPGKDSKGNSKRAET*
                                                                  (SEQ ID NO: 426)
       9 PL0123
                                : EQVGGLQPGRGTPGKDSKGNAKRAET+
                                                                  (SEQ ID NO: 427)
     10 HV3F$HUMAN
                               : EQVGGLQPGRGTPGKDSKGDSRRAET+
                                                                  (SEQ ID NO: 428)
      11 JL0048
                                : EQVGGLQPGRGTPGKDSKGNSRRAET+
                                                                  (SEQ ID NO: 429)
     12 HV3B$HUMAN
                                : QQVGGLEPGRGTPGKDSKGBSKRAET*
                                                                  (SEQ ID NO: 430)
     13 HUMIGHBV
                                : EQLGDLQPGRGTPGKASKGNSKRAET+
                                                                  (SEQ ID NO: 431)
     14 HV3E$HUMAN
                                : EQVGGLQPGRGTTGKDSKGDSKRAET+
                                                                  (SEQ ID NO: 432)
     15 PL0116
                                : QQVGGVQPGRGTPGKDSKGNSKRAET*
                                                                  (SEQ ID NO: 433)
     16 HV3K$HUMAN
                                : QQVGGVQPGRGIPGKDSKGNSKRPET*
20
                                                                  (SEQ ID NO: 434)
     17 N$2PB4H
                                : EQVGGVQPGRGIPGKDSKGDSKRPET*
                                                                  (SEQ ID NO: 435)
     18 HV3I$HUMAN
                                :QQVGGVQPGRGTPGKDSHGDSKRPET*
                                                                  (SEQ ID NO: 436)
     19 HV3J$HUMAN
                               :QKVGGVQPGRGTPGKDSKGNSKRTET*
                                                                  (SEQ ID NO: 437)
     20 HV3G$HUMAN
                               :QEVGGVZPGRGTPGKBSKGBSKRAET*
                                                                  (SEQ ID NO: 438)
     21 HV3M$HUMAN
                                : EQLGGLQPGRGTPGKDSNGDSKQAZT+
                                                                  (SEQ ID NO: 439)
                               : EQLGGLQPGRGSPGKDTNGDSKEA2T+
     22 HV3O$HUMAN
                                                                  (SEQ ID NO: 440)
25
     23 HV3N$HUMAN
                                : AQLGGLQPGRGTPGKDSNGDSKQAZS+
                                                                  (SEQ ID NO: 441)
     24 HV3R$HUMAN
                               : EQLGGLQPGRGTPGKVSQGDSKQAZT*
                                                                  (SEQ ID NO: 442)
     25 HV3P$HUMAN
                               : EQVGGLQPGRGTPGKVSQGDSKEP2T*
                                                                  (SEQ ID NO: 443)
     26 HUMIGHCV
                               : EQLGGLQPERGTPGKESKGHSMRAET+
                                                                  (SEQ ID NO: 444)
     27 HV3TSHUMAN
                               : EQVGDLQPGRGBPGKDSKGNAKRVET+
                                                                  (SEQ ID NO: 445)
     28 HV3U$HUMAN
                               : EQVGDLQPGRGNPGKDSKGNAQRPET+
                                                                  (SEQ ID NO: 446)
30
     29 PL0098
                               : QQVGGVQPGRGTLGKDSKGNSKRAET+
                                                                  (SEQ ID NO: 447)
     30 HV3H$HUMAN
                               :QZVGGAZPGRGSPGKASKGBSKRAET+
                                                                  (SEQ ID NO: 448)
     31 HV3ASHUMAN
                               : QQVGGLKPGRGSPGKDSKGNAQRTZT*
                                                                  (SEQ ID NO: 449)
     32 HV3SSHUNAN
                               : DQVGGLKPGRGTPGKHSHGDSKTP2T*
                                                                  (SEQ ID NO: 450)
     33 HUMIGHAN
                               : EOLGGLOPGRGTSREDSKGNSKRAET*
                                                                  (SEQ ID NO: 451)
     34 HV3Q$HUKAM
                               : Eqvgalqpgrgtpgkdsqadskeazt*
                                                                  (SEQ ID NO: 452)
35
     35 A36040
                               : EQLGGLQPGRGTPGK----VEGSVET*
                                                                  (SEQ ID NO: 453)
     36 HUMIGHAM
                               : EQVGAFQPGRGNSGKASKGDSKRPDT+
                                                                  (SEQ ID NO: 454)
                                                                  (SEQ ID NO: 455)
     37 HUMIGHAO
                               : EQVGAPQPGKGNSGKASKGDSKRPDT+
     38 HUMIGHAR
                               : EQVGAFQPGKGNSGKASKGDSNRPDT*
                                                                  (SEQ ID NO: 456)
     39 HV3L$HUMAN
                               :QQVGGVQAGRAMPGKDSRGISKRTET*
                                                                  (SEQ ID NO: 457)
                                                                  (SEQ ID NO: 458)
     40 HVLASHUMAN
                               :QQVAEVKPGKGTPGQQKQGESTRSET+
     41 A32483
                               :QQVAEVKPGKGTPGQQKQGTSTRSET*
                                                                  (SEQ ID NO: 459)
40
                               :QQVAEVKPGKGTPGQQKQGTSARSET+
     42 HUMIGHAY
                                                                  (SEQ ID NO: 460)
     43 HUMIGHCU
                               : QQVAEVKPGKGTPGQQKQGTSIRSDT*
                                                                  (SEQ ID NO: 461)
     44 HUNIGHBS
                               : QQVAEVKPGKGTPGQEKQGTSIRSDT*
                                                                  (SEQ ID NO: 462)
     45 HUNIGVHLS
                               :QQVAEVKPGKGTPGQQHQGTSTRSDT*
                                                                  (SEQ ID NO: 463)
     46 HUNIGHBX
                               :QQVGEVKPGRGTPGQQKQDTSTRSDT+
                                                                  (SEQ ID NO: 464)
                               : QQVAEVKPGRGTPGHPRQGASFRSDS*
     47 HV1C$HUMAN
                                                                  (SEQ ID NO: 465)
45
     48 H34964
                               :QQVSELKPGKGTPGQQGTGTSVKAET*
                                                                  (SEQ ID NO: 466)
                               : Eqvaevkpgkgspgkpsqgksikast*
     49 HUMIGHCY
                                                                  (SEQ ID NO: 467)
                               : eqvaevkpgrgspgkpsqgksikast*
                                                                  (SEQ ID NO: 468)
     50 PL0119
```

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			(SEQ	ID I	NO:	4691
	51 HV1F\$HUMAN	: QQVAEVKPGRGDPGRPRQASSTISAT*	(SEQ			470)
5	52 D34964	: EQVAEVPQGKGRPGKSLQGKSLKAST*				471)
		: QQMAEVKPGRGTPGKPGVVPSFFSET*	,		NO:	•
	• • • • • • • • • • • • • • • • • • • •	:QQVAEVKPGRGTPGRYIWEPSFFNEG*	(SEQ		NO:	472)
	54 HV1E\$HUMAN	:QQQAGLKPSSGSPGKPSKSTSKTAAT*	(SEQ	ID	NO:	473)
	55 JL0047	:QQQPGLKPSSGSPGKPSKSTSKTAAT*	(SEQ	ID	NO:	474)
	56 HUMIGHBW	COOPGLEPSSOSPORTSKSTSWTAAT	(SEQ	מז	NO:	475)
	57 E34964	: QQQPGLKPSSGSPGKPSKSTSNTAAT*			NO:	476)
10	58 HUMIGHCW	:QQQPGLKPSSGSAGKPSKSTSKTAAT*			NO:	477)
	59 HV2FSHUMAN	:ROOPGLKPSSGPPGKPSRGTSRSAAT*	, -			
		:OOOAGLKPSSGSPGRTSKSTSKTAAT*	, -		NO:	478)
		: OOPPGLRPSSGTPGRTPRSTSKTAAT*	(SEQ		ΝО:	479)
	61 HV2G\$HUMAN	: XQEPGLRPSSGSPGRTPRSTSKTAAT*	(SEQ	ID	NO:	480)
	62 N\$3FABH	:QQQPGLKPSSGSPSRVSKSTSKTPET*	(SEQ	ID	NO:	481)
	63 PS0091	:QHQAGLKRSSGPPGKPSTSTSKTAAT*		ID	NO:	482)
15	64 HUMIGHDA	OHOAGLARSGPPORPSIGNOR	(SEQ	ID	NO:	483)
15	65 A26555	: ZQESGLKPTSGSPGKPSKSRSKAADA*	,			484)
	66 HV2ESHUMAN	:QTKPTLKPTTGSPGRPSKSTSKDPVT*	(SEQ			•
		:OTKPTLKPTTGSPGKPSRSTSRDPVS*	(SEQ	ID	NO:	485)
	<u> </u>	· PTRPALKPTTGSPGKTSKTTSKDPVT*	(SEQ	ID	ио:	
	68 A36005	:QNRPALKATTGSPGKTSETTSKDPAT*	(SEQ	ID	NO:	487)
	69 HV2H\$HUHAN	:OTTPALKPRTGSPGKTSRTDSKNPVT*	(SEQ	ID	NO:	488)
	70 HV2ASHUMAN	:QTRPALRPTTGSPGEASETTSKGPGT*	(SEQ			
20	71 HV2C\$HUMAN	:UIRPALKFI 103F0EMBETTSEDTAY*	(SEQ			
	72 HV2B\$HUMAN	:QTRPALKPTTGSPGKTSETTSRDTAY*	•		NO:	:
	73 JL0049	:LEGVQLWGGRGISRKYAKGNGKRDES*	(SEQ	τD	NO.	73+1
	/3 484443					

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EXAMPLE 2

DETAILED DESCRIPTION OF METHOD FOR CONSTRUCTING THREE-DIMENSIONAL MODEL OF ANTIBODY VARIABLE REGION

The references cited in the text below are listed at the end of this Example.

The first antibody Fab structure was determined in 1972. Since then, no more than about twelve Fab structures have been published, a number that represents a very small fraction of the total antibody repertoire (>108 antibodies). To understand the molecular basis of this antibody diversity will require knowledge of either a large number of x-ray structures, or the rules by which combining site topography is governed. The development of such prediction rules has now reached the point where variable regions of antibodies can be modelled to an accuracy approaching that of the medium resolution x-ray structure.

The interaction of an antibody with its cognate antigen is one of the most widely accepted paradigms of molecular recognition. To understand the antibody-antigen interaction in atomic detail requires knowledge of the three-dimensional structure of antibodies and of their antigen complexes. Traditionally such information has come from x-ray crystallographic studies (see Davies et al. for review (Davies et al., 1988)).

The modelling of antibody combining sites was first attempted by Padlan & Davies (Padlan et al., 1976) at a time when very few antibody structures were known. Nonetheless, Padlan and colleagues recognized that the key lay in high structural homology that existed within the β-sheet framework regions of different antibody variable domains. The antigen combining site is formed by the juxtaposition of six interstrand loops, or CDRs (Complementarity Determining Regions) (Kabat et al., 1987), on this framework. If the framework could be modelled by homology then it might be possible to model the CDRs in the same way. Padlan and Davies (Padlan et al., 1976) reasoned that CDR length was the important determinant of backbone conformation though the number of antibody structures was insufficient to thoroughly test this maximum overlap procedure (MOP). This notion was not picked up again until the early 1980's when Pedersen and Rees proposed a similar approach to modelling antibody combining sites based on a more extensive analysis of antibody structures (de la Pas et al., 1986).

Those essentially knowledge-based procedures are best exemplified for antibodies by the work of Chothia & Lesk (Chothia et al., 1986) who, in 1986, extended and modified the MOP procedure by introducing the concept of "key" residues. These residues allow the further subdivision of CDRs of the same length into "canonical" structures which differ in having residues at specified positions that, through packing, hydrogen bonding or the ability to assume unusual values of the torsion angels ϕ , ψ and ω , determine the precise CDR conformation

(Chothia et al., 1989). Similar knowledge-based methods have been proposed for predicting loop conformations in general (Thornton et al., 1988; Tramontano et al., 1989). These methods rely on the crystallographic database of protein structures. However, none of the above knowledge-based methods has been totally successful. In particular, the MOP or canonical structure approaches have succeeded in modelling only five of the six CDRs. This stems from the fact that the third CDR of the heavy chain, H3, is more variable in sequence, length and structure than any of the other CDRs.

To deal with this problem several groups have attempted to use ab initio methods to model the combining site (Bruccoleri and Karplus, 1987). The requirement with such methods is that the total allowable conformational space accessible to a particular CDR is sampled. Typical of purely geometric approaches is that of Go & Sheraga (Go and Sheraga, 1970) and more recently Palmer & Sheraga (Palmer and Sheraga, 1991), where the problem is reduced to one in which the central region of the polypeptide backbone, having characteristic bond length and bond angles, is constructed between the end points of the loop (CDR if an antibody loop) by a "chain closure" algorithm. In a modification of this algorithm, Bruccoleri & Karplus (Bruccoleri and Karplus, 1987) introduced an energy minimization procedure which greatly expanded the domain of conformational space searched during the chain closure procedure. This modification is incorporated into the conformational search program CONGEN (Bruccoleri and Karplus, 1987), which also allows the user to choose any set of standard bond length and bond angels such as the CHARMM (Brooks et al., 1983) standard geometry parameter sets. Other approaches such as minimization (Moult and James, 1986), or molecular dynamics (Fine et al., 1986) either fail to saturate conformational space or are unable to deal with the problem of long CDRs. Whichever of the ab initio methods is employed however, the problem is one of defining the selection criteria in such a way as to allow the unambiguous identification of the correct structure (in this context correct is defined by reference to an appropriate X-ray structure) within the ensemble of candidates, for every CDR. To date this has not been possible.

Recently a more holistic approach has been taken to the modelling of CDRs which combines the advantages of knowledge-based and *ab initio* methods in a single algorithm known as CAMAL (Combined Algorithm for Modelling Antibody Loops) (Martin et al., 1989; Martin et al., 1991). Previously this algorithm has been used to model individual CDRs in the presence of the crystal structure conformations of the other five. As is demonstrated below, CAMAL is able to predict the backbone conformations of all six CDRs of the antibody combining site to an accuracy approaching that of medium resolution x-ray structures. In addition the algorithm includes a procedure for selecting and fitting together the light and heavy chain framework regions prior to generation of CDR conformations, thus making possible the prediction of the entire variable region. Furthermore a new Monte Carlo (MC) simulated annealing method has been developed for the determination of sidechain conformations.

35 The Framework Region

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Antibody framework regions consist of conserved β -strands that form the β -barrel structure characteristic of immunoglobulin V-type regions. In the procedure described here each V-region is built from a database of known antibody structures, using sequence homology for selection of the light (L) and heavy (H) chain V-domains. The two domains are then paired by least squares fitting on the most conserved strands of the antibody β -barrel (Table 2 and Figures 5 & 6. The strand orientations were determined by analyzing the barrels of known antibody crystal structures. Eight antibodies were analyzed using a multiple structure fitting program as follows. Seven structures were fitted onto one of the set selected at random and mean coordinates were calculated. All eight structures were then fitted onto these mean coordinates and new mean coordinates determined. This procedure was iterated until the mean coordinate set converged (5-10 cycles). The variance for the mean coordinates at each barrel point (N,C α ,C) was calculated. In Figure 5 this variance is plotted against the projected positions of these points onto the conjugate axis of the barrel.

Strand 8 and all but two residues of strand 7 in both light and heavy chains were eliminated as they showed deviations greater than 3 σ (standard deviation units) from the mean coordinates. These two strands comprised the takeoff points of CDR H3, and suggests that any knowledge-based prediction of CDR H3 would have to account not only for sequence and length variation in the CDR itself, but also for the position of the participating strands. The remaining mean coordinates were used as a scaffold onto which the L and H chains were fitted. Strands 7 and 8 in the final framework were obtained from the database structure used in the construction. The framework strands are marked + in the multialignment in Table 2.

The sidechains were then replaced using a 'maximum overlap' method, in which sidechain templates were fitted on backbone atoms with the sidechain torsion angles being adjusted to match those of equivalent torsions in the parent sidechain.

The Combining Site

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The procedure for predicting the structure of combining sites combines a database search with a conformational search procedure. The architecture of the program suite to perform this task is outlined in Figure 7.

The database search utilizes distance constraints for each of the six CDR loops determined from known antibody structures. These constraints were determined by calculating $C\alpha$ - $C\alpha$ distances within known loops and using a search range of $x+3.5\sigma$ (the mean ± 3.5 standard deviation units). A database containing all the proteins in the Brookhaven Protein Databank (Bernstein et al., 1977) is then searched for fragments which satisfy the constraints for a loop of the required length. The middle section of the loop is then deleted and reconstructed using the conformational search program CONGEN (Bruccoleri and Karplus, 1987). For loops of six or seven residues, the structure database appears to saturate the conformational space available to the backbone adequately and only sidechains are built by conformational search. Loops shorter than six residues are built by conformational search alone since this is computationally feasible and the number of loops selected from the database becomes unacceptably large as loop length decreases.

When modelling a complete combining site, loops of 6 or more residues are modelled individually with the other loops absent. If the loops are built consecutively, small errors can accumulate leading to a poor result (Martin, 1990). All the loop conformations are then evaluated using a solvent modified potential, which excludes the attractive van der Waals and electrostatic terms of the non-bonded energy function contained within the GROMOS (Aqvist et al., 1985) potential. The lowest five energy conformations are selected and filtered using a "structurally determining residue" algorithm (FILTER), based on backbone torsion angles observed in the original database loops. Since the database search is not used for the shortest loops of 5 residues or fewer, the FILTER algorithm cannot be used. Energy is thus the only available selection criterion and the short loops are built last, in the presence of the longer loops.

Side Chains

The determination of sidechain positions was previously done using the iterative sidechain determination algorithm described by Bruccoleri et al. (Bruccoleri and Karplus, 1987). Unfortunately the CHARMM (Brooks et al., 1983) force field fails to select the correct conformations of exposed hydrophobic sidechains. There is no penalty for having an exposed uncharged atom, without solvent present. CONGEN is also unable to saturate the conformational space for a large number of sidechains (more than 6 residues).

Recently Lee et al. (Lee and Levitt, 1991; Lee and Subbiah, 1991) has proposed a method for searching conformational space for a large number of sidechains using MC simulated annealing. A simple energy function is used for the evaluation of conformations generated by a biased random walk:

$$E = \sum_{i=1}^{n} \epsilon_{o} \left(\left(\frac{r_{o}}{r} \right)^{6} - 2 \left(\frac{r_{o}}{r} \right)^{12} \right) + \kappa_{o} \cdot COS \left(3 \omega \right)$$

Where the first term is a simple Lennard-Jones potential which evaluates the non-bonded contacts between the atoms in a given molecule, the second term is a simple torsional term which only applies to C-C bonds. The torsional term biases the function towards 60° rotamers. ε_o and κ_o are constants. The metropolis function: $P = C \frac{-\delta E}{T}$

$$P = C^{-\delta E}$$

is used to evaluate the energy function. Any move which results in a decrease in energy is accepted, and any move which results in a positive δE is only accepted with the probability P. This simple method can be used to search the large conformational space defined by a set of torsion angles in amino-acid sidechains, and find or define the global minimum which exist for a set of sidechains. T is the simulation temperature.

When searching sidechain conformations using this method the simulation system usually gets trapped in an energetic minima well before the global minimum is encountered, at a high temperature, without the solution space having been searched sufficiently. This problem can be solved by truncating the Lennard-Jones potential, thus allowing atoms to pass through each other. In reality this function would converge towards infinity when the distance r between the atoms approaches zero.

The evaluation of sidechain conformations generated is done solely on the basis of energy, for internal (core) residues, since good van der Waal's interactions are considered to be equal to a good packing of the sidechains. The situation becomes more complicated when trying to predict the conformation of surface residues. The lowest van der Waal's interaction is obtained by a combination of sidechain conformations which minimize the overlap of atoms, this means that the lowest energy is obtained with extended conformations of sidechains, without considering good packing of sidechains.

Using the fact that hydrophobic, bulky residues will be shielded by the hydrophilic sidechains, and will be buried in the surface, it is possible to generate a simple function which will evaluate these macroscopic observations. These functions can either be implemented in the objective evaluation function of the Monte Carlo simulation, or as is done here, added as a post processing step. Including an accessibility/hydrophobicity term in the evaluation function would slow down the calculation considerably, hence the term has been added as a post processing function. The function used is a sum of the product of relative exposed surface area multiplied by the residual hydrophobicities. The hydrophobicities used are taken from Cornette et al. (Cornette et al., 1987).

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$$f_{conformation} = \sum_{i=1}^{n} -A_{irel} \cdot H_{irel}$$

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n is the number of sidechains reconstructed. The surface area is calculated using the tesselated icosahedron approach (Chau and Dean, 1987), which is not very precise (0.1 percent), but is able to evaluate a large number of conformations. The function is evaluated for the final 2,000 conformations and the lowest value conformation selected as the best.

Using this simple approach it is possible to integrate over a large phase space with many degrees of freedom, and get a complete sampling of the space.

Predicted Structures of an Anti-hapten, Anti-peptide and Two Anti-protein Antibodies

In the following section the predicted structures of four different antibody F_V regions are presented and analyzed. The antibodies are:

- Gloop-2 (Darsley and Rees, 1985), an anti-lysozyme antibody whose Fab structure was determined by Jeffrey et al., (Jeffrey et al., 1991) and which was used as a learning exercise during the development of CAMAL.
- D1.3 (Amit et al., 1986), an anti-lysozyme antibody whose uncomplexed F_V coordinates were supplied by R. Poljak et al. after the model coordinates had been deposited.
- 36-71 (Rose et al., 1990), an anti-phenylarsonate antibody whose Fab structure was carried out by D.
 R. Rose, et al., and whose coordinates were obtained after the model coordinates had been deposited.
- 3D6 (Grunow et al., 1988), an anti-protein (GP41 of HIV) antibody whose Fab structure was carried out by D. Carter et al. (Carter, 1991) and whose coordinates were obtained after the model coordinates had been deposited. For this antibody, the model was generated using the canonical loop method of Chothia & Lesk (Chothia et al., 1989; Chothia et al., 1986) for CDRs L1, L2, H1 and H2, while L3 and H3, which cannot be modelled using canonical structures, were constructed using CAMAL.

All four models were subjected to both restrained and unrestrained energy minimization using the DIS-COVER (TM Biosym Technology) potential with 300 cycles of steepest descents, followed by conjugate gradient minimization until convergence to within 0.01 Kcal occurred.

The resolution and R-factors of the x-ray structures are given in Table 3 together with the parent frameworks selected in building the models. The structures and models were compared by global fits of the loops. The β -barrel strands 1 to 6, as described above, were least squares fitted and the RMS deviation was then calculated over the loops. The backbone (N,C α ,C) RMS values for fitting model and crystal structure frameworks were between 0.4 and 0.9 Å, illustrating the conservation of the core β -barrel. Using all eight strands RMS deviations between 0.6 and 1.2 Å were observed.

Global fits (Table 4) give a more realistic measure of the accuracy of the model than a local least-squares fit over the loops since they account for the overall positioning of the loops in the context of the F_V structure. Local fits, which give lower RMS deviations, are also shown in Table 4. Differences between local and global RMS deviations arise from differences in V_H/V_L domain packing and differences in loop 'take off' angles and positions.

Table 5 shows the canonical loops selected from modelling 3D6. Backbone structures of the modelled CDRs, superimposed on the x-ray structures after global fitting are shown in Figure 8. General features and points of interest for each of the six CDRs are discussed below.

Analysis of the CDR Regions

During the comparison of CDR conformations in the V-region models and the x-ray Fab structures it was observed that at certain positions in a CDR, the peptide backbone may adopt either of two conformations by undergoing a "peptide flip" (1,4 shift). This phenomenon is also seen in type 2 β -turns (Paul et al., 1990). Dynamics simulations of β -turns show that the transformation energy between $\phi 1 = -00$, $\psi 1 = -30$, $\phi 2 = -90$, $\psi 2 = 0$ and $\phi 1 = -00$, $\psi 1 = 120$, $\phi 2 = 90$, $\psi 2 = 0$ has a maximum value of 5 kcal (Paul et al., 1990). This is low enough to allow selection of either conformation. The peptide flip is observed within several canonical classes (as described by Chothia et al. (Chothia et al., 1989)) and the hydrogen bonding pattern used to determine the conformation of a canonical class does not disallow the peptide flip. Any modelling procedure should therefore take these, or any other multiple conformations, into consideration where the transformation energies are sufficiently low to permit population of the different conformational forms. Table 6 shows an example of the "peptide-flip" phenomenon from the crystallographic database of antibody structures. It should be noted that a single crystal structure will not show multiple conformations since the crystallization will 'freeze out' one of the conformations. During the modelling procedure the two populations of conformers are easily extracted from a set of *ab initio* generated loops, by using a torsional clustering algorithm.

CDR-L1

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In Gloop-2 and D1.3, all five low energy conformations were very similar with RMS deviations differing by less than 0.25 Å (backbone) and 0.35 Å (all atoms). The FILTER algorithm was unable to distinguish between the conformations and the lowest energy structure was selected.

Although CDR-L1 of 3D6 was originally built using the canonical loop from HyHEL-10, the mid-section was rebuilt by conformational search, for the following reason. HyHEL-10 and REI CDR-L1 loops are placed in the same canonical ensemble (Chothia et al., 1989) although they contain a 1-4 shift (peptide flip) relative to one another between the fifth and eighth residues of the loop (residues 28-31) (see Table 6).

36-71 shows the same 1-4 shift between the model and crystal structure CDRs. Both crystal structure and model were compared with other loops of the same canonical class as defined by Chothia et al. (Chothia et al., 1989). It was found that the hydrogen bonding pattern which determines the conformation was conserved.

CDR-L2

CDR-L2 of D1.3 has two adjacent threonines (49, 50) which in the x-ray structure are packed against the tyrosine at the fourth position of CDR-H3, thus minimizing the exposed hydrophobic sidechains. In the unminimized model the threonine sidechains are exposed to the solvent, but after energy minimization, this packing is observed.

CDR-L3

In Gloop-2, D1.3 and 36-71 the proline at the seventh position in the loop is correctly predicted in the *cis* conformation. It has previously been suggested that the conformation of CDR-L3 is dictated by the presence of a proline in position 8 or 9 (Chothia et al., 1989) within the loop. 3D6 does not have a proline in either position. Only 7 out of 290 CDR-L3 sequences (Kabat et al., 1987) lack a proline at both positions and in all of the published x-ray structures this proline is present. This is an example of a situation where either a new canonical class may need to be defined or where the canonical rule breaks down altogether, and an alternative method must be employed.

The 3D6 L3 loop is 7 residues in length and was built using database loops alone where conformational space is saturated by means of fragments selected from the crystallographic database (Global RMS: 2.01 Å, N,C α ,C), and by using CAMAL (Construction: Q[Q(YNS)Y]S, Global RMS: 1.97 Å, N,C α ,C). The similarity of the structures generated by the two procedures illustrates the utility of the database search and suggests that, for shorter loops it is capable of saturating the available conformational space.

CDR-H1

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Using the Kabat and Wu definition of CDR-H1 places this loop as an extension of the β -sheet. The extended nature of this stretch of peptide limits its conformational flexibility and CDR-H1 is generally modelled accurately (Martin et al., 1989; Chothia et al., 1989).

In Gloop-2 and D1.3, the Phe or Tyr sidechain at the second position in the loop is poorly placed and packs against Leu at the penultimate position in HFR1 (see Table 2). 36-71 has a well-placed Asn at this position, rather than the more common bulky hydrophobic sidechain.

CDR-H2

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CDR-H2 of 36-71 is similar in sequence to F19.9 (Strong et al., 1991), (36-71: YNNPGNGYIA (SEQ ID NO:492); F19.9: YINPGKGYLS (SEQ ID NO:493)). While the structurally determining residues specified by Chothia and Lesk (Chothia et al., 1989) are conserved, the backbone conformations are different: F19.9 has a bulge at the -PGN- Gly, compared with 36-71, giving the loop a 'kink' in the middle. The model of 36-71 shows a 1-4 shift, though the sidechains are still well placed.

In Gloop-2, the all atom RMS deviation is poor (3.00 Å) (Jeffrey et al., 1991) when compared with the P2₁ crystal structure, owing to rotations of the Phe at position 3 in the loop and Tyr at position 10 by approximately 120° about the χ_2 torsion angle. Gloop-2 has been solved in two different crystal forms, P2₁ and P1 (Jeffrey et al., 1991; Jeffrey, 1989). When compared with the P1 structure, the sidechains are placed almost perfectly and the all atom RMS (global fit) drops to 2.23 Å.

This concerted sidechain motion between crystal forms illustrates the effects of crystallization conditions on surface sidechain placement. Even though surface sidechains may show low temperature factors indicating low mobility in the crystal, their mobility in solution may be high. In the Gloop-2 P1 structure, the mean sidechain temperature factor for the F_V domain is 13.46 (σ = 8.20) while the sidechains of these two residues of H2 show mean temperature factors of 5.56 (σ = 0.68) for the Phe at position 3 and 7.10 (σ = 1.73) for the Tyr at position 10.

CDR-H3

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CDR-H3 is the most variable of the six CDR's with all lengths up to 21 residues being represented in Kabat et al., (Kabat et al., 1987). This extreme variability results from V-D-J splicing (Schilling et al., 1980) and has always been a problem when attempting to model antibodies. Such loops may be divided into short (up to 7 residues), medium (up to 14 residues) and long (15 or more residues). Using the CAMAL procedure, short and medium CDR-H3's can be modelled as accurately as other CDR's of similar lengths. Although long CDR-H3's are more difficult and cannot, at present, be built to the same accuracy, the chain trace is still correct.

It is unlikely that the longer loops consist of 'pure' loops (i.e., all random coil or turn). In crystal structures of antibodies with medium to long CDR-H3 loops (McPC603 (Rudikoff et al., 1981): 11 amino acids (aa); KOL (Marquart et al., 1980): 17 aa; F19.9 (Lascombe et al., 1989): 15 aa) the loops consist of a disordered β -sheet extension from the β -barrel core and a 5-8 residue random coil/turn connecting these two strands.

To determine the nature of medium to long loops (>8 residues) which satisfy the CDR-H3 constraints, a complete search of the Protein Databank for loops of length 8-20 residues, was performed using the inter-C α distance constraints determined from known antibody crystal structures for CDR-H3. The resulting loops were then analyzed using the DSSP (Kabsch and Sander, 1983) program, which is able to assign secondary structure to polypeptide structures. The amount of secondary structure for each length of loop was calculated, and it was observed that for loops longer than 12 residues the amount of secondary structure within each of the classes described in DSSP was constant. The number of loops selected is also constant (approximately 150 loops) for loops longer than 12 residues. A closer inspection of each of the length ensembles shows indeed that the loops are the same between the groups.

This analysis shows that, like the long CDR-H3 crystal structures, the selected fragments consist of β -strands connected by 5-8 residue loops. For loops above 12-13 residues in length, the same loops are selected, but with extensions to the β -strands. This is called the "sliding-ladder" effect. In addition, the maximum size of a random coil or turn fragment in any of the structures contained in the Protein Databank tends not to exceed 8 residues, as determined by DSSP. This implies that the conformational space of longer loops is not saturated by the database and, although it is unlikely that long loops in antibodies will differ significantly from long loops in other structures, confidence in the prediction must be correspondingly reduced.

By how much is the usefulness of the CAMAL algorithm reduced by this observation?

The frequency of occurrence of different CDR-H3 lengths in antibody sequences described by Kabat et al. (Kabat et al., 1987) was analyzed. Figure 10 shows that more than 85% of H3 loops have lengths between 4 and 14 residues which can be modelled accurately by the CAMAL algorithm.

CDR-H3 of D1.3 is of average length (8 residues), though no loops of this length are seen in the available antibody structures. The crystal structure coordinate set showed an RMS of 1.9 Å compared with the model. The 36-71 loop is 12 residues long. The conformation is correctly predicted as a short loop connecting an

extension of the β -sheet.

The 3D6 H3 loop is 17 residues long. While KOL (Marquart et al., 1980) has the same length it has only one residue in common with 3D6 and only one conservative mutation. There is thus no reason to believe that the conformations would be similar. The final predicted conformation of 3D6 is an extended β -sheet, as in the crystal structure. The difference between the predicted and the crystal structure of 3D6-H3 is due to a twist of 5-7° in the extended β -sheet loop (see Figures 9A-9D). Such a twist has also been observed for complexed and uncomplexed antibodies by Wilson et al. (Wilson and others). This suggests that long CDR-H3 loops may be flexible and actively involved in antigen binding.

10 The Complete Variable Region

Prediction of the strand positions and $V_L - V_H$ orientation in the framework β -barrel was exact for all of the four antibodies. The backbone (N,C α ,C) RMS deviations from the crystal structures were between 0.56 and 0.86 Å, despite the fact that, in all cases the V_L and V_H regions of a particular model were derived from different antibody structures. This suggests that this method will do well in procedures such as humanization (Gorman et al., 1991), where correct framework positioning is important. The backbones of all six CDRs in all four antibodies are essentially correctly predicted, as shown in Figure 8. There are two important points to make about these predictions. First, the position of each CDR on its framework barrel is correct. Thus, CDR-framework interactions can be confidently monitored. The only deviation from the x-ray structure is CDR-H3 of antibody 3D6 which has been previously discussed. Second, the all atom RMS deviation between models and x-ray structures is dominated by sidechain positions. In most instances this deviation is due to a small number of incorrectly positioned, exposed sidechains (for example, in D1.3 the only sidechains which are incorrectly predicted are Tyr 9 of L1, Trp 4 of L3, Tyr 2 of H1 and Tyr 4 of H3). Since each CDR is constructed in the absence of other CDRs, the force field may choose a rotamer which is 120° away from that found in the crystal structure. This effect has also been observed by Lee et al. (Lee and Levitt, 1991).

Conclusion

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For antibodies having CDR H3 regions of 14 residues or less the complete variable domain can be modelled to an accuracy approaching that of medium resolution x-ray structures. For antibodies with longer H3 loops the CAMAL algorithm is likely to need an additional procedure in which molecular dynamics simulations are also incorporated.

The canonical approach of Chothia et al. appears to work well (at least in modelling backbones) where it may be applied and may be used successfully in combination with the CAMAL procedure.

One important observation that has emerged from these studies is that a given loop can exist in several conformations. In particular, this seems likely for CDR-L1 and, to a lesser extent, CDR-L3 and longer CDR-H3's. A simple combinatorial calculation shows that, if each of these three loops can exist in three separate conformations, a given combining site can have $3^3 = 27$ different topographies. Clearly, this would explain the origins of cross reactivity and would allow for induced fit of antigens.

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10	ent of ar d region: L - Cha chain be	- 000 - +222 + 0 + 0 + 0 + 0 + 0 + 0 + 0 + 0 +	D - QMTQS C
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20	Table 2: Alignment of antibody sequences used in the modelling. '*' indicates CDR, regions; '+' indicates β -strand regions used in the fitting for modelling frameworks. Nomenclature for β -barrel strands is (H or L - Chain) - FR(Framework region)-(Strand number), thus for example strand one of the heavy chain becomes HFR1.		
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4 5	"*" indicates CDR, regions; '+' rks. Nomenclature for β -barrel ber), thus for example strand	FFFF	++QQQQ+ ++TTT++ ++QQQQ+ ++TTT++ ++QQQQ+ ++TTT++ ++QQQQ+ ++QQQQ+ ++QQQQ+ ++QQQQ+ ++QQQQ+ ++QQQQ+
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			Framew	ork Model
Antibody	Resolution	R-factor	Light	Heavy
Gloop-2	2.80	21.2	REI	HyHEL-5
D1.3	-	-	REI	NEW
36-71	1.90	20.9	Gloop2	NEW
3D6	2.70	17.7	REI	KOL

Table 3: Details of the antibody crystal structures against which the models were compared and the parent frameworks used to build the models. Resolution data for D1.3 has not yet been published.

	_						2				NA N		
op-3 L1 RASIG(EIS)GIYLS 484 0.71 3.08 1.94 0.87 3.08 3 RASIG(NIE)NIFLA 484 3.29 1.83 4.80 3.81 3.43 4.88 3.81 3.41 3.48 3.88 3.81 3.41 3.48 3.40 4.89 3.21 3.43 4.80 3.81 3.41 3.48 3.40 4.89 3.21 3.41 3.40 3.40 3.41 3.41 3.41 3.40 3.40 3.41 3.41 3.41 3.41 3.40 3.40 3.41		Astibody	CDR	90 G S 0 3 C 0	SEQ ID NO	3	N.C.	VII CG	All MC	Ce		All Co	VII MC
Depth Company of the													
RASIG(DIN)NIFLN 493 2.29 1.29 4.34 4.34 3.94 2.72 3.43 4.34 4.35 4.36 4.36 4.37 4.36 4.37 4.37 4.37 4.37 4.38		Gloop-2	1.1	RASIQ(EIS)GIYLS	•	0.73	0.71	2.01	-	0.0	0.87	3.9	2.13
RASIQ(SIO)N NLH 496 2.71 2.43 2.40 2.80 3.81		DI.3		RASIG(NIH)NIYLA	3	2.20	1.93	4.34	9.94	2.73	2.48	4.50	4.82
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AASTLDS AASTLDS APPLICATION APPLICAT	_	3D6		RASIO(SIG)NINCH	197	0.81	0.54	2.48	1.02	0.81	0.70	2.00	-
AASTLDS AAASTLDS AAASTLDS AAASTLDS AAASTLDS AAAA AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA	_								•		}	: :	:
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KASSLES SOL O.41 O.42 1.37 1.20 O.48 O.48 O.49		36-71		FIT(SRS)QIS	500	0.64	0.64	2.34	2.28	0.73	0.73	2.41	2.40
Op-2 L3 LQ[Y(LSY)P]LT 503 0.58 0.58 1.73 1.60 0.76 0.76 2.00 3 Q4[G(NAL)P]RT 503 1.41 1.38 2.60 2.98 1.76 1.79 3.46 71 Q[Q(YNS)Y]S 504 1.00 1.00 2.26 2.10 1.40 1.30 2.97 71 [T(FGI)T] 506 0.40 0.70 2.00 2.31 1.01 2.04 9p-2 H1 [T(FGI)Y] 506 0.44 0.42 2.33 2.00 0.43 0.90 3.24 9p-2 H2 EI[F(PGN)S]KTY 510 0.43 0.44 0.42 2.33 1.10 0.90 3.24 9p-2 H2 EI[F(PGN)S]KTY 510 0.43 0.47 1.53 1.10 0.81 0.90 3.24 11 VNN[PGN]HTA 511 0.42 0.42 0.43 1.53 1.10 0.81 0.90 3.24 11 ISWDSSIG 513 0.43 0.52 2.33 1.00 0.81 1.01 1.02 11 ER[D(YRL)D]Y 514 0.50 0.53 0.43 3.43 3.00 0.81 <		3D6		KASSLES	801	0.41	0.43	1.37	1.30	0.83	0.00	1.73	1.80
OP-2 LG[F(KST)P]FL			•		<u> </u>	• •	3	:	5	9 7 8	0.74	8	
71 QQ[Q(NNS)Y]S 504 1.00 1.00 2.36 3.10 1.40 1.30 3.87 0p-2 H1 [T(FGI)T] 506 0.60 0.70 3.00 1.60 3.00 3.00 3.01 1.01 2.04 0p-2 H1 [S(NGI)N] 500 0.60 0.70 0.83 2.33 2.00 0.48 0.90 3.24 0p-2 H2 EI[F(PQN)S]KTY 510 0.60 0.63 2.23 1.11 0.61 0.72 2.81 0p-2 H2 EI[F(PQN)S]KTY 510 0.63 0.64 1.63 1.70 1.20 0.61 0.72 2.81 0p-2 H2 EI[F(PQN)S]KTY 511 0.62 0.42 0.63 1.60 1.04 0.97 2.61 1 YNNIP(GDG)N TD 511 0.42 0.43 0.62 1.55 1.40 0.67 0.64 1.60 1 ER[GYRDY]DI 514 0.64 0.65 2.35 2.05 0.64 1.60 1.60 0.65 0.65 0.6			5		502	1.41	1.95	2.80	2.92	1.76	1.79	:	3.30
T(FGI)T SOG 0.40 0.70 2.00 1.00 1.01 2.04 3.70 3.60		36-71		OOIG(NAL)PIRT	504	-0	T.00	2.24	3.10	1.1	1.34	2.87	2.20
Treatity		3D6		QQ(YNS)YIS	308	1.40	1.88	3.04	3.90	2.81	1.97	2	3.3
T(FGI)T SOC COLOR COL	_							}	}	:	:		3
G(YGV)N S07 0.44 0.62 2.38 2.00 0.83 0.77 0.24 0.97 0.24 0.97 0.24 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25 0.97 0.25		Gloop-3	Ξ	[T(FGI)T]	506	0.00	0.70	2.00	-				
Second Nation Second S	_	D1.3		[O(YGV)N]	507	0.44	0.63	2.33	2.00	0.85	0.90	3.24	2.5
DYAMH SIP (PGN)S KTY SIQ 0.43 0.43 1.43 1.70 1.20 0.84 2.23 1.86 1.40 0.87 0.86 1.40 0.87 0.86 1.40 0.87 0.86 1.40 0.87 0.88 1.40 0.80 0.80 0.80 0.80 0.80 0.80 0.80 0		36-71		(S(NQI)N)	500	0.00	0.84	2.22		1.04	0.97	2.51	2.23
DP-2 H2 EI[F(PGN)S]KTY 510 0.03 0.04 1.03 1.70 1.20 0.94 2.23 MI[W(GDG)N]TD 511 0.02 0.42 1.55 1.40 0.97 0.95 1.00 1.00 1.00 1.00 1.00 1.00 1.00 1.0	_	3D6		DYAMH	509	0.47	0.77	1.83	1.11	0.81	0.73	1.69	1.20
MI[W(GDG)N]TD MI[W(GDG)N]TD VNN[P(GNG)Y IA 110 110 110 110 110 110 110 1		3	5	EIIF/PONISIKTY	5		•	<u>.</u>	1.70	- 2	0.94	2.28	2.10
P-2 H3 [R(E)R)Y] S14 0.66 0.59 2.68 2.09 0.67 1.07 2.68 2.09 0.67 1.07 2.68 2.09 0.67 1.07 2.68 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 0.69 2.69 2.69 2.69 2.69 2.69 2.69 2.69 2		D1.3	;	MIW(GDG)NITD	51.	3	0.43	1.54	T. 68	0.87	0.88	-	1.86
	_	36-71		YNN(P(GNG)Y IA	513	0.0	0.78	2.01	J.30	1.47	1.61	1.7	-
P-2 H3 [R(EIR)Y]	_	3D6		ISWDSSSIG	513	0.48	0.52	2.88	2.08	0.95	0.89	2.86	2.10
ER(D(YFID)Y 515 0.30 0.83 1.00 1.20 0.01 1.90 1.90 1.90 1.90 1.90 1.90 1.9			<u> </u>			5 2 2		9	*	0. 0	1.07	.	• •
SEYY[O(GSY)K]FDY 516 1.98 1.75 4.40 6.00 2.68 2.83 4.40 6.30 3.96 5.30	_	21.3	;	EBCD(VBC)DIV	515 515	0 5	0.53	1.00	2	1.25	2.2	<u>.</u>	1.33
GRDYYID/SGG)YFITVAFDI 517 3.66 3.42 8.93 4.01 4.30 3.96 6.30		36-71		SEYYIQIQSYIKIPDY	316	1.95	1.75	4.6	<u>•</u>	2.65	2.53	8	.0
	_	3D6		GRDYY(D(SGG)YF)TVAFDI	517	3.66	3.42	5.93	4.01	4.80	3.96	9.30	9.30

0.86 and 0.56 respectivly calculating the RMS over the loops. The total RMS of the frameworks (N,C α ,C) is 0.81, 0.60, calculated by least-squares fitting the conserved core of the two structures upon each other and difference between model and crystal structure loop coordinates. The RMS values are a global fit Table 4: Sequence and conformational search construction scheme for each of the 24 CDRs, =construction area, ()= Chain closure, all sidechains are constructed. RMS(Root Mean Square)

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Loop	Canonical	Sequence	SEQ ID NO
Ll	HyHEL-10	RASQSISRWLA	518
	(3D6)	RASQSIGNNLH	497
L2	REI	EASNDLA	519
	(3D6)	KASSLES	501
H1	McPC603	DFYME	520
	(3D6)	DYAMH	509
H2	KOL	IIWDDGSDQ	521
	(3D6)	ISWDSSSIG	513

Table 5: Canonical loops selected for the model of 3D6(taken from Chothia et al (1989)).

Residu	e Number	24	25	26	27	28*	29*
REI	Sequence	Q	Α	S	Q	S	I
ILDI	ϕ/ψ	-/138	-103/157	-96/7	-158/142	-40/108	-112/9
HvHEL-10	Sequence	R	Á	Ś	Q	S	I
Hynt DD-10	ϕ/ψ	-/108	-85/135	-88/64	172/160	-64/-38	9/63
Resid	ie Number	30*	31* .	32	33	32	
REI	Sequence	I	K	. Y	L	N	SEQ ID NO: 52
1422	ϕ/ψ	79/-77	-146/21	-104/89	-143/133	-144/-	
HyHEL-10	Sequence	G	N	Ń	Ĺ	H	SEQ ID NO: 51
Hyneb-10	ϕ/ψ	-63/107	85/-15	-105/72	-129/118	-126/-	

Table 6: Backbone ϕ and ψ angles of residues in CDR-L1 from HyHEL-10 and REI classified in the same canonical group by Chothia *et al* (1989). The residues exhibiting a peptide flip are indicated by a *.

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Oxford, UK. 5 SEQUENCE LISTING GENERAL INFORMATION 10 (i) APPLICANT: PEDERSEN, Jan T. SEARLE, Stephen M.J. Anthony R. REES, ROGUSKA, Michael A. GUILD, Braydon C. 15 (ii) TITLE OF INVENTION: SURFACE RESIDUE VENEERING OF RODENT ANTIBODIES (iii) NUMBER OF SEQUENCES: 522 20 (iv) CORRESPONDENCE ADDRESS: (A) ADDRESSEE: Sughrue, Mion, Zinn, Macpeak & Seas (B) STREET: 2100 Pensylvania Avenue, N.W. (C) CITY: Washington (D) STATE: D.C. 25 (E) COUNTRY: United States (F) ZIP: 20037-3202 (v) COMPUTER READABLE FORM: (A) MEDIUM TYPE: Floppy disk (B) COMPUTER: HP 9000/700 Workstation 30 (C) OPERATING SYSTEM: UNIX (D) SOFTWARE: In house (vi) CURRENT APPLICATION DATA: (A) APPLICATION NUMBER: 07/942,245 35 (B) FILING DATE: 09-SEP-1992 (C) CLASSIFICATION: (ix) TELECOMMUNICATION INFORMATION: (A) TELEPHONE: (202) 293-7060 40 (B) TELEFAX: (202) 293-7860 (C) TELEX: 6491103 (1) INFORMATION FOR SEQ ID NO:1 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 109 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1: Asp Ile Gln Met Thr Gln Ser Pro Ser Ser Leu Ser Ala Ser Leu Gly 15 10 5 55

5	Glu	Arg	Val	Ser 20	Leu	Thr	Cys	Arg	Ala 2		Gln	Glu	Ile	_	Gly 30	Tyr
10	Leu	Ser	Trp 35	Leu	Gln	Gln	Lys	Pro 40	Asp)	Gly	Thr	Ile		Arg 5	Leu	Ile
	Tyr	Ala 50	Ala	Ser	Thr	Leu	Asp 55	Ser	Gly	Val	Pro		Arg O	Phe	Ser	Gly
15	Arg 65	Arg	Ser	Gly	Ser	Asp 70	Tyr	Ser	Leu	Thr	Ile 75	Ser	Ser	Leu	Glu	Ser 80
	Glu	Asp	Phe	Ala	Asp 85	Tyr	Tyr	Cys	Leu	Gln 90		Leu	Ser	Tyr	_	Leu 95
20	Thr	Phe	Gly	Ala 100	Gly	Thr	Lys	Leu	Glu 105		ı Lys	s Ar	y Ala	à		
	(2)	INFO	RMAT:	ION	FOR	SEQ	ID N	0:2								
25			(i) :	(1	A) L B) T	ENGT YPE:	RACT H: 1 ami OGY:	09 a no a	mino cid		.ds					٠.
30				MOLE SEQUI			_	_		Q ID	NO:	2:		-		
35	Asp 1	Ile	Gln	Met	Thr 5		Ser	Pro	Ala	Ser 1		Ser	Ala	Ser		Gly L5
	Glu	Thr	Val	Thr 20	Ile	Thr	Cys	Arg	Ala 25		Gly	Asn	Ile	_	Asn O	Tyr
40 -	Leu	Ala_	Trp 35	Tyr	Gln	Gln	Lys	Gln 40		Lys	Ser	Pro	_	Leu 5	Leu	Val
45	Tyr	Tyr 50	Thr	Thr	Thr	Leu	Ala 55		Gly	Val	Pro	_	Arg O	Phe	Ser	Gly
50	Ser 65	Gly	Ser	Gly	Thr	Gln 70	Tyr	Ser	Leu	Lys	Ile 75	Asn	Ser	Leu	Gln	Pro 80
	Glu	Asp	Phe	Gly	Ser	Tyr	Tvr	Cvs	Gln	His	Phe	Trn	Ser	Thr	Pro	Arq

5	Thr Phe Gly Gly Thr Lys Leu Glu Ile Lys Arg Arg 100 105
	(3) INFORMATION FOR SEQ ID NO:3
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 107 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:
	Asp Ile Val Leu Thr Gln Ser Pro Ala Ile Met Ser Ala Ser Pro Gly 1 5 10 15
20	Glu Lys Val Thr Met Thr Cys Ser Ala Ser Ser Ser Val Asn Tyr Met 20 25 30
25	Tyr Trp Tyr Gln Gln Lys Ser Gly Thr Ser Pro Lys Arg Trp Ile Tyr 35 40 45
	Asp Thr Ser Lys Leu Ala Ser Gly Val Pro Val Arg Phe Ser Gly Ser 50 55 60
30	Gly Ser Gly Thr Ser Tyr Ser Leu Thr Ile Ser Ser Met Glu Thr Glu 65 70 75 80
35	Asp Ala Ala Glu Tyr Tyr Cys Gln Gln Trp Gly Arg Asn Pro Thr Phe 85 90 95
	Gly Gly Gly Thr Lys Leu Glu Ile Lys Arg Ala 100 105
40	(4) INFORMATION FOR SEQ ID NO:4
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 109 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
_. 45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

Asp Ile Val Leu Thr Gln Ser Pro Ala Thr Leu Ser Val Thr Pro Gly 1 5 10 15

5	Asn	Ser	Val	Ser 20	Leu	Ser	Cys	Arg	Ala 25		Gln	Ser	Ile	_	Asn O	Asn
	Leu	His	Trp 35	Tyr	Gln	Gln	Lys	Ser 40		Glu	Ser	Pro		Leu 5	Leu	Ile
	Lys	Tyr 50	Ala	Ser	Gln	Ser	Ile 55		Gly	Ile	Pro	Ser 6		Phe	Ser	Gly
15	Ser 65	Gly	Ser	Gly	Thr	Asp 70	Phe	Thr	Leu	Ser	Ile 75	Asn	Ser	Val	Glu	Thr 80
	Glu	Asp	Phe	Gly	Met 85	Tyr	Phe	Сув	Gln	Gln 9	_	Asn	Ser	Trp		Tyr 5
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	Gln	Lys	Val	Thr 20		Thr	Cys	Ser	Ala 2		Ser	Ser	Val		Ser 30.	Leu
40	His	Trp	Tyr - 35		Gln	Lys	Ser		Thr 0	Ser	Pro	Lys	Pro	Trp 45	Ile	Tyr
45	Glu	Ile 50	ser	Lys	Leu	Ala	Ser 5		Val	Pro	Ala		Phe	Ser	Gly	Ser
50	Gly 65		Gly	Thr	Ser	Tyr 70		Lev	Thr	: Ile	* Asr 75	Thr	Met	Glu	Ala	Glu 80
	Asp	Ala	Ala	Ile	Tyr 89		Cys	Glr	Glr	Trp	Thr	Tyr	Pro	Leu	lle	Thr 95

Phe Gly Ala Gly Thr Lys Leu Glu Leu Lys Arg Ala

(6) INFORMATION FOR SEQ ID NO:6

10	(A) LENGTH: 112 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:
	Glu Ser Val Leu Thr Gln Pro Pro Ser Ala Ser Gly Thr Pro Gly Gln 1 5 10 15
20	Arg Val Thr Ile Ser Cys Thr Gly Thr Ser Ser Asn Ile Gly Ser Ile 20 25 30
25	Thr Val Asn Trp Tyr Gln Gln Leu Pro Gly Met Ala Pro Lys Leu Leu 35 40 45
30	Ile Tyr Arg Asp Ala Met Arg Pro Ser Gly Val Pro Thr Arg Phe Ser 50 55 60
	Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Glu 65 70 75 80
35	Ala Glu Asp Glu Ser Asp Tyr Tyr Cys Ala Ser Trp Asn Ser Ser Asp 85 90 95
	Asn Ser Tyr Val Phe Gly Thr Gly Thr Lys Val Thr Val Leu Gly Gln 100 105 110
4 0	(7) INFORMATION FOR SEQ ID NO:7
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 115 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:
	Asp Ile Val Met Thr Gln Ser Pro Ser Ser Leu Ser Val Ser Ala Gly 1 5 10 15
55	

5	Glu	Arg	Val	Thr 20	Met	Ser	Cys	Lys	Ser 2		Gln	Ser	Leu	_	Asn 0	Ser
	Gly	Asn	Gln 35	Lys	Asn	Phe	Leu	Ala 40		Tyr	Gln	Gln		Pro 5	Gly	Gln
10	Pro	Pro 50	Lys	Leu	Leu	Ile	Tyr 55		Ala	Ser	Thr	_	Glu O	Ser	Gly	Val
15	Pro 65	Asp	Arg	Phe	Thr	Gly 70	Ser	Gly	Ser	Gly	Thr 75	Asp	Phe	Thr	Leu	Thr 80
	Ile	Ser	Ser	Val	Gln 85	Ala	Glu	Asp	Leu	Ala 90		Tyr	Tyr	Cys		Asn 5
20	Asp	His	Ser	Tyr 100	Pro	Leu	Thr	Phe	Gly 10		Gly	Thr	Lys	Leu 11		Ile
25	Lys	Arg	Ala 115									•	,			
	(8)	INFO	RMAT:	ION	FOR	SEQ	ID N	8:0								
30			(i) :	· (.	ENCE A) L B) T C) T	ENGT YPE:	H: 1 ami	.03 a	mino		ds					
		(.	ii) 1	MOLE	CULE	TYP	E: p	epti	.de							
35		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	NO:	8:				
	Ser 1	Val	Leu	Thr	Gln 5		Pro	Ser	Val	Ser 1		Ala	Pro	Gly		Arg L5
40	Val	Thr_	Ile	Ser 20	Cys	Thr	Gly	Ser	Ser 2			Ile	_	_	Gly 0	Asn
45	His	Val	Lys 35	_	Tyr	Gln	Gln	Leu 40		Gly	Thr	Ala		Lys 5	Leu	Leu
50	Ile	Phe 50	His	Asn	Asn	Ala	Arg 55	_	Ser	Val	Ser		Ser 0	Gly	Ser	Ser
•	Ala 65		Leu	Ala	Ile	Thr 70	Gly	Leu	Gln	Ala	Glu 75	Asp	Glu	Ala	Asp	Tyr 80
55																

Tyr Cys Gln Ser Tyr Asp Arg Ser Leu Arg Val Phe Gly Gly Thr 85 90 95

	Lys Leu Thr Val Leu Arg Gln 100
10	(9) INFORMATION FOR SEQ ID NO:9
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 114 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:
20	Asp Val Val Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly 1 5 10 15
25	Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Leu Val His Ser 20 25 30
	Gln Gly Asn Thr Tyr Leu Arg Trp Tyr Leu Gln Lys Pro Gly Gln Ser 35 40 45
30	Pro Lys Val Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly Val Pro 50 55 60
35	Asn Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile 65 . 70 . 75 . 80
•	Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Phe Cys Ser Gln Ser 85 90 95
40	Thr His_Val Pro Trp Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys 100 105 110
45	Arg Ala
	(10) INFORMATION FOR SEQ ID NO:10
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 109 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55	(ii) MOLECULE TYPE: peptide

		(:	X1)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	NO:	10:				
5	Asp 1	Ile	Gln	Met	Thr 5	Gln	Thr	Thr	Ser	Ser 1	Leu 0	Ser	Ala	Ser		Gly IS
10	Asp	Arg	Val	Thr 20	Ile	Ser	Cys	Arg	Ala 25		Gln	Asp	Ile	_	Asn 0	Tyr
15	Leu	Asn	Trp 35	Tyr	Gln	Gln	Lys	Pro 40		Gly	Thr	Val		Leu 5	Leu	Val
	Tyr	Tyr 50	Thr	Ser	Arg	Leu	His 55		Gly	Val	Pro	_	Arg O	Phe	Ser	Gly
20	65			Gly		70					75					80
25	Glu	Asp	Ile	Ala	Thr 85	Tyr	Phe	Cys	Gln	Gln 90	_	Ser	Thr	Thr	_	Arg 5
				Gly 100		-			105		Lys	Arq	y Arq	J		
30	(11)			SEQU! (;) (1)	ENCE A) L: B) T	CHA ENGT YPE:		ERIS 09 a no a	TICS mino cid		.ds					
35		•		MOLE SEQUI			_	-		Q II	NO:	11:				
40	Asp 1	Ile	Gln	Met	Thr 5		Ile	Pro	Ser	Ser		Ser	Ala	Ser		Gly .5
15	Asp	Arg	Val	Ser 20	Ile	Ser	Cys	Arg	Ala 25		Gln	Asp	Ile		Asn O	Phe
45	Leu	Asn	Trp 35	Tyr	Gln	Gln	Lys	Pro 40		Gly	Thr	Ile		Leu 5	Leu	Ile
50	Tyr	Phe 50	Thr	Ser	Arg	Ser	Gln 55		Gly	Val	Pro		Arg 0	Phe	Ser	Gly

5	Ser 65	Gly	Ser	Gly	Thr	Asp 70	Tyr	Ser	Leu	Thr	11e ·75	Ser	Asn	Leu	Glu	Gln 80
•	Glu	Asp	Ile	Ala	Thr 85	Tyr	Phe	Cys	Gln	Gln 90	_	Asn	Ala	Leu		Arg 5
10	Thr	Phe	Gly	Gly 100	Gly	Thr	Lys	Leu	Glu 105		Lys	Arç	g Ala	1		
•	(12)	INF	ORMA'	TION	FOR	SEQ	ID	NO:1	2							
15			(i)	(I	A) L B) T	ENGT YPE:		07 a	minc	aci	ds		•			
20		(ii)	MOLE	CULE	TYP	E: p	epti	.de					-		
20		(xi)	SEQU	ENCE	DES	CRIF	MOIT	: SE	EQ II	NO:	12:				
25	Asp 1	Ile	Gln	Met	Thr 5		Ser	Pro	Ser	Thr 1		Ser	Ala	Ser		Gly L5
20	Asp	Arg	Val	Thr 20		Thr	Cys	Arg	Ala 2		Gln	Ser	Ile		Arg 30	Trp
30	Leu	Ala	Trp 35	Tyr	Gln	Gln	Lys	Pro		Lys	Val	Pro		Leu 15	Leu	Ile
35	Tyr	Lys 50		Ser	Ser	Leu	Glu 5		Gly	Val	Pro		Arg 50	Phe	Ser	Gly
,	Ser 65	_	Ser	Gly	Thr	Glu 70		Thr	Leu	Thr	Ile 75		Ser	Leu	Gln	Pro 80
40	Asp	Asp	Phe	Ala	Thr 85		Tyr	Cys	Gln		Tyr o	Asn	Ser	Tyr	Ser	Phe 95
45	Gly	Pro	Gly	Thr 100		s Vai	l As _l	o Il	e Ly: 10		g Th	r				
45	(13)	INF	FORMA	MOITA	FOE	R SE	Q IĎ	NO:	13							
50			(i)		(A) I (B) I	LENG' FYPE	ARAC' TH: : am LOGY	104 ino	amin acid	o ac	ids					
			(ii)	MOLI	CUL	Е ТҮ	PE:	pept	ide							

		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:
5	Gln 1	Val Gln Leu Gln Gln Ser Gly Thr Glu Leu Ala Arg Pro Gly Ala 5 10 15
10	Ser	Val Arg Leu Ser Cys Lys Ala Ser Gly Tyr Thr Phe Thr Thr Phe 20 25 30
;	Gly	Ile Thr Trp Val Lys Gln Arg Thr Gly Gln Gly Leu Glu Trp Ile 35 40 45
15	Gly	Glu Ile Phe Pro Gly Asn Ser Lys Thr Tyr Tyr Ala Glu Arg Phe 50 60
20	Lys 65	Gly Lys Ala Thr Leu Thr Ala Asp Lys Ser Ser Thr Thr Ala Tyr 70 75 80
	Met	Gln Leu Ser Ser Leu Thr Ser Glu Asp Ser Ala Val Tyr Phe Cys 85 90 95
25	Ala	Arg Glu Ile Arg Tyr Trp Gly
	(14)	INFORMATION FOR SEQ ID NO:14
30		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 107 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:
40	Gln 1	Val Gln Leu Lys Glu Ser Gly Pro Gly Leu Val Ala Pro Ser Gln 5 10 15
	Ser	Leu Ser Ile Thr Cys Thr Val Ser Gly Phe Ser Leu Thr Gly Tyr 20 25 30
45	Gly	Val Asn Trp Val Arg Gln Pro Pro Gly Lys Gly Leu Glu Trp Leu 35 40 45
50	Gly	Met Ile Trp Gly Asp Gly Asn Thr Asp Tyr Asn Ser Ala Leu Lys 50 55 60

5	Ser 65	Arg	Leu	Ser	Ile	Ser 70	Lys	Asp	Asn	Ser	Lys 75	Ser	Gln	Val	Phe	Leu 80
	Lys	Met	Asn	Ser	Leu 85	His	Thr	Asp	Asp	Thr 90		Arg	Tyr	Tyr		Ala 5
10	Arg	Glu	Arg	Asp 100	Tyr	Arg	Leu	Asp	Tyr 105		Gly	7 .				
	(15)	INFO	ORMA'	TION	FOR	SEQ	ID	NO:1	.5							
15		1	(i)	Ċ	A) L B) T	ENGT YPE:	H: 1 ami		mino cid		ids					
		(:	ii)	MOLE	CULE	TYF	E: p	epti	de							
20		(:	xi)	SEQU	ENCE	DES	CRIE	OIT	I: SI	EQ II	ои с	:15:				
	Val	Gln	Leu	Gln	Gln 5		Gly	Ala	Glu		Met 0	Lys	Pro	Gly		Ser 15
25	Val	Lys	Ile	Ser 20		Lys	Ala	Ser	Gly 2		Thr	Phe	Ser	Asp	Tyr 30	Trp
30	Ile	Glu	Trp 35		Lys	Gln	Arg	Pro 4		His	Gly	Leu	Glu	Trp 15	Ile	Gly
35	Glu	Ile 50		Pro	Gly	Ser	Gly 5		Thr	Asn	Tyr	His	Glu 50	Arg	Phe	Lys
	Gly 65	Lys	Ala	Thr	Phe	Thr		Asp	Thr	Ser	Ser 75	Ser	Thr	Ala	Tyr	Met 80
40	Gln	Leu	Asn	Ser	Leu 8		Ser	Glv) Asp	Ser 9	Gly	Val	. Tyr	Tyr	Cys	Leu 95
45	His	Gly	Ası	ту: 100		Ph	e As	p Gl	y Tr 10		Y .					
45	(16)	INF	ORM	ATIO	1 FO	R SE	Q ID	NO:	16							
50	٠		(i)		(A) : (B) :	LENG TYPE	TH: : am	104 ino	STIC amin acid near	o ac l	ids					
			(ii)	MOT	• /											

		(:	XI)	SEQU.	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	16:				
	Asp 1	Val	Gln	Leu	Gln 5		Ser	Gly	Pro	Ser		Val	Lys	Pro		Gln .5
10	Thr	Leu	Ser	Leu 20	Thr	Cys	Ser	Val	Thr 25		Asp	Ser	Ile		Ser 0	Asp
15	Tyr	Trp	Ser 35	Trp	Ile	Arg	Lys	Phe 40		Gly	Asn	Arg		Glu 5	Tyr	Met
	Gly	Tyr 50	Val	Ser	Tyr	Ser	Gly 55		Thr	Tyr	Tyr	Asn 6	_	Ser	Leu	Lys
20	Ser 65	Arg	Ile	Ser	Ile	Thr 70	Arg	Asp	Thr	Ser	Lys 75	Asn	Gln	Tyr	Tyr	Leu 80
	Asp	Leu	Asn	Ser	Val 85	Thr	Thr	Glu	Asp	Thr 90		Thr	Tyr	Tyr		Ala 5
25 [°]	Asn	Trp	Asp	Gly 100	Asp	Tyr	Trp	Gly								
30	(17)			(1	ENCE A) L B) T	CHA ENGT YPE:	RACT H: 1 ami	ERIS	TICS mino cid		.ds					
35		ţ		MOLE			_	•		. TE	NO.	. 17.				
40	Glu 1	. •	•	SEQU!		Glu				-	Leu		Gln	Pro		Gly .5
	Ser	Leu	Lys	Leu 20	Ser	Cys	Ala	Ala	Ser 25		Phe	Asp	Phe		Lys 0	Tyr
45	Trp	Met	Ser 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Ile
50	Gly	Glu 50	Ile	His	Pro	Asp	Ser 55		Thr	Ile	Asn		Thr 0	Pro	Ser	Leu

5	Lys 65	Asp	Lys	Phe	Ile	Ile 70	Ser	Arg	Asp	Asn	Ala 75	Lys	Asn	Ser	Leu	Tyr 80
	Leu	Gln	Met	Ser	Lys 85	Val	Arg	Ser	Glu	Asp 90		Ala	Leu	Tyr	Tyr 9	Cys 5
10	Ala	Arg	Leu	His 100	Tyr	Tyr	Gly	Tyr	Asn 105		Туг	Tr	Gly	Z ,		
	(18)	INF	ORMA!	TION	FOR	SEQ	ID	NO:1	8							
15			(i) :	(1	A) L: B) T	ENGT YPE:		17 a no a	minc	: aci	.ds					
20		(:	ii)	MOLE	CULE	TYP	E: p	epti	de							
20		(:	xi)	SEQU:	ENCE	DES	CRIP	TION	: SE	Q II	NO:	:18:				
25	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1	_	Val	Gln	Pro	Gly 1	Arg .5
	Ser	Leu	Arg	Leu 20		Cys	Ser	Ser	Ser 2		Phe	Ile	Phe		Ser 30	туr
30	Ala	Met	Tyr 35	_	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 15	Trp	Val
35	Ala	Ile 50		Trp	Asp	Asp	Gly 5		Asp	Gln	His		Ala 50	Asp	Ser	Val
	Lys 65		Arg	Phe	Thr	Ile 70	Ser	Arg	Asn	Asp	Ser .75	Lys	Asn	Thr	Leu	Phe 80
40	Leu	Gln	Met	Asp	Ser 85		Arg	Pro	Glu	Asp 9		Gly	Val	Tyr	Phe	Cys 95
45	Ala	Arg	Asp	Gly 100		His	Gly	Phe	Cys 10		Ser	Ala	Ser	Cys 1	Phe 10	Gly
	Pro	Asp	Tyr 115	Trp	Gly	•		•								
50	(19)	INF	ORMA	TION	FOF	SEÇ	Q ID	NO:	19							
			(i)		JENCE (A) I (B) I	LENG?	rh:	113	amin	s: o ac	ids					
55				ľ	"											

				1	C) 1	OPOL	OGY:	110	ear							
5		(ii)	MOLE	CULE	TYP	E: p	epti	de							
		(xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	NO:	19:				
10	Glu 1	Val	Lys	Leu	Val 5		Ser	Gly	Gly	Gly 1		Val	Gln	Pro	_	G1) L5
	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Thr	Ser 25		Phe	Thr	Phe	_	Asp 0	Phe
15	Tyr	Met	Glu 35	Trp	Val	Arg	Gln	Pro		Gly	Lys	Arg		Glu 5	Trp	Ile
20	Ala	Ala 50	Ser	Arg	Asn	Lys	Gly 55		Lys	Tyr	Thr		Glu O	Tyr	Ser	Ala
25	Ser 65	Val	Lys	Gly	Arg	Phe 70	Ile	Val	Ser	Arg	Asp 75	Thr	Ser	Gln	Ser	Ile 80
	Leu	Tyr	Leu	Gln	Met 85	Asn	Ala	Leu	Arg	Ala 9		Asp	Thr	Ala		Tyr 95
30	Tyr	Сув	Ala	Arg 100		Tyr	Tyr	Gly	Ser 10		Trp	Tyr	Phe	Asp 11	_	Trp
35	Gly											•				•
	(20)	INF	ORMA	TION	FOR	SEQ	ID	NO: 2	0							
40	-		(i) : 	(A) L B) T	ENGT YPE:	H: 1 ami		mino cid	S: o aci	ids					
•		(ii)	MOLE	CULE	TYP	E: p	epti	de		*	,				
45		(xi)	SEQU	ENCE	DES	CRIF	TION	: SI	II QE	ONO:	20:				
	Val 1	Gln	Leu	Glu	Gln 5		Gly	Pro	Gly	Leu 1	_	Arg	Pro	Ser		Thi 15
50	Leu	Ser	Leu	Thr 20		Thr	Val	Ser	Gly 2		Ser	Phe	Asp		Tyr 30	Туз

5	Ser	1111	35	vai	Arg	GIII	Pro	Pro 40		Arg	СТĀ	Leu		Trp 5	Ile	Gly
	Tyr	Val 50	Phe	Tyr	His	Gly	Thr 55	Ser	Asp	Thr	Asp		Pro 0	Leu	Arg	Ser
10	Arg 65	Val	Thr	Met	Leu	Val 70	Asn	Thr	Ser	Lys	Asn 75	Gln	Phe	Ser	Leu	Arg 80
15	Leu	Ser	Ser	Val	Thr 85	Ala	Ala	Asp	Thr	Ala 90		Tyr	Tyr	Cys	_	Arg 95
	Asn	Leu	Ile	Ala 100	Gly	Cys	Ile	Asp	Val 105	_	Gly	7				
20	(21)	INFO	RMA	rion	FOR	SEQ	ID	NO:2	1 .							
		(i) S	(1	A) L:	ENGT:	H: 1 ami	ERIS 09 a no a lin	mino cid		.ds					
25		(i	 i) N		•			epti								
							_	TION		o to	NO:	21:				
30	Glu 1	Val									Leu		Gln	Pro	_	Arg .5
35	Pro	Met :	Lys	Leu 20	Ser	Cys	Val	Ala	Ser 25		Phe	Thr	Phe	_	Asp 0	Tyr
	Trp	Met .	Asn 35	Trp	Val	Arg	Gln	Ser 40		Glu	Lys	Gly		Glu 5	Trp	Val
40	Ala	Gln_ 50	Ile	Arg	Asn	Lys	Pro 55		Asn	Tyr	Glu	Thr 6		Tyr	Ser	Asp
45	Ser 65	Vál :	Lys	Gly	Arg	Phe 70	Thr	Ile	Ser	Arg	Asp 75	Asp	Ser	Lys	Ser	Ser 80
	Val	Tyr	Leu	Gln	Met 85	Asn	Asn	Leu	Arg	Val 90		Asp	Met	Gly		Tyr 5
50	Tyr	Cys	Thr	Gly 100	Ser	Tyr	Tyr	Gly	Met 105	_	Туг	Tr	Gly	Y		,
55	(22)	INFO	RMAT	MOI	FOR	SEQ	ID	NO:2	2							

5			(1)	(1	A) L B) T	ENGT YPE:	RACT H: 1 ami OGY:	15 a no a	mino cid		ds					
10				MOLE SEQU			_	-		Q II	NO:	22:				
	Gln 1	Val	Gln	Leu	Lys 5		Ser	Gly	Ala	Glu 1		Val	Ala	Ala		Ser 5
15	Ser	Val	Lys	Met 20	Ser	Cys	Lys	Ala	Ser 25		Tyr	Thr	Phe		Ser 0	Tyr
20	Gly	Val	Asn 35	Trp	Val	Lys	Gln	Arg 40		Gly	Gln	Gly		Glu 5	Trp	Ile
25	Gly	Туг 50	Ile	Asn	Pro	Gly	Lys 55		Tyr	Leu	Ser	_	Asn O	Glu	Lys	Phe
	Lys 65	Gly	Lys	Thr	Thr	Leu 70	Thr	Val	Asp	Arg	Ser 75	Ser	Ser	Thr	Ala	Tyr 80
30	Met	Gln	Leu	Arg	Ser 85	Leu	Thr	Ser	Glu	Asp 9		Ala	Val	Tyr		Cys 5
35 ·	Ala	Arg	Ser	Phe 100	Tyr	Gly	Gly	Ser	Asp 10	-	Ala	Val	Tyr	Tyr 11	_	Asp
	Ser	Trp	Gly 115													
40	(23)			TION SEQU						: :						
			-	Ċ	в) т	YPE:	H: 1 ami OGY:	no a	cid	aci	ids					
45		·	•	MOLE			_	_								
	Gl 11	•	-	SEQU Leu									Ara	Ala	Glv	Ser
50	1	•			5			1		1	-		9			15

5	Ser	Val	Lys	Met 20	Ser	Cys	Lys	Ala	Ser 25		Tyr	Thr	Pne	Thr 3		Asn
10	Gly	Ile	Asn 35	Trp	Val	Lys	Gln	Arg 40		Gly	Gln	Gly	Leu 4		Trp :	Ile
	Gly	Tyr 50	Asn	Asn	Pro	Gly	Asn 55		Tyr	Ile	Ala	Tyr 6		Glu	Lys	Phe
15	Lys 65	Gly	Lys	Thr	Thr	Leu 70	Thr	Val	Asp	Lys	Ser 75	Ser	Ser	Thr	Ala	Tyr 80
20	Met	Gln	Leu	Arg	Ser 85		Thr	Ser	Glu	Asp 90		Ala	Val	Týr		Cys 5
	Ala	Arg	Ser	Glu 100		Tyr	Gly	Gly	Ser 10		Lys	Phe	Asp	Tyr 11	Trp .0	Gly
25	(24)									٠						
30			(i)	(A) I B) T	CHA ENGT YPE: OPOL	H: :	l17 a ino a	amino acid	aci	ids					·.
		•	•	MOLE						PO TI	n NO	. 24 .				
35	Glu	•	-	SEQU Leu									Gln	Pro	Gly	Arg
	1		,			5		_		1	.0					15
40	Ser	Leu	Arg	Leu 20		Cys	Ala	Ala	Ser 2	Gly	Phe	Thr	Phe	Asn	Asp 30	Tyr
	Ala	Met	His 35		Val	. Arg	Glr	n Ala 4	Pro	Gly	Lys	Gly	Leu	Glu 45	Trp	Val
45	Ser	Gly 50	_	e Ser	Trp) Asp		ser 55	: Ser	: Ile	Gly	Tyr	Ala 60	. Asp	Ser	Val
50	Lys 65		/ Arc	g Phe	e Thi	70		r Arq	j Asp) Asr	n Ala 75	Lys	. Asr	n Ser	Leu	Tyr 80
	Leu	Gli	n Met	t Asr	n Sei 8		ı Ar	g Ala	a Glu	ı Asr	Met 90	: Ala	. Lev	1 Туг	Tyr	Cys 95

5	. Val	Lys	Gly	Arg 100	Asp	Tyr	Tyr	Asp	Ser 105		Gly	Tyr	Phe	Thr 11		Ala
	Phe	Ąsp	Ile 115	Trp	Gly											
10	(25)	INF	ORMA	TION	FOR	SEQ	ID	NO:2	5							
15			(i) :	()	ENCE A) L B) T C) T	ENGT YPE:	H: 1	11 a no a	minc		.ds					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
		(:	xi) a	SEQU:	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	25:				
20	Asp 1	Val	Leu	Met	Thr 5		Thr	Pro	Leu	Ser 1		Pro	Val	Ser		Gly L5
25	Asp	Gln	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25	_	Gln	Ile	Ile	_	His 0	Ser
30	Asp	Gly	Asn 35	Thr	Tyr	Leu	Glu	Trp		Leu	Gln	Lys		Gly 5	Gln	Ser
50	Pro	Lys 50	Leu	Leu	Ile	Tyr	Lys 55	_	Ser	Asn	Arg	_	Ser 0	Gly	Val	Pro
35	Asp 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Asp 75	Phe	Thr	Leu	Met	Ile 80
40	Ser	Arg	Val	Glu	Ala 85		Asp	Leu	Gly	Val		Tyr	Сув	Phe		Gly 95
	Ser	His	_Val	Pro 100	His	Thr	Phe	Gly	=	y Gly		_		u Gl:	_	e
45	(26)	INF	orma	TION	FOR	SEQ	ID	NO: 2	6							
			(i)	(ENCE A) L B) T C) T	ENGT	H: 1 ami	.10 a	mind		ids					
50		(ii)	MOLE	CULE	TYF	E: p	epti	de							
		(xi)	SEQU	ENCE	DES	CRIE	PTION	1: SI	EQ II	ои с	:26:				

5	Gln 1	Ser	Val	Leu	Thr 5	Gln	Pro	Pro	Ser	Ala 1		Gly	Thr	Pro		Gln L5
10	Arg	Val	Thr	Ile 20	Ser	Cys	Ser	Gly	Thr 25		Ser	Asn	Ile		Ser 0	Ser
	Thr	Val	Asn 35	Trp	Tyr	Gln	Gln	Leu 40		Gly	Met	Ala	_	Lys 5	Leu	Leu
15	Ile	Tyr 50	Arg	Asp	Ala	Met	Arg 55		Ser	Gly	Val		Asp 0	Arg	Phe	Ser
20	65			Ser		70					75				-	80
	Ser	Glu	Asp	Glu	Thr 85	Asp	Tyr	Tyr	Cys	Ala 9		Trp	Asp	Val		Leu 5
25				Val 100					105	_	va]	Thi	. Vai	l Lei 110	_	
	(27)	INF	ORMA'	TION	FOR	SEQ	ID	NO:2	7							
30				(1	A) L: B) T' C) T(ENGT YPE: OPOL	H: 1 ami OGY:	ll a no a lin	mino cid ear		.ds					
35		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
•		(:	xi) :	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	NO:	27:				
40	Gln 1	Val	Leu	Met	Thr 5		Thr	Pro	Ser	Ser 1	_	Pro	Val	Thr		Gly .5
	Gln	Gln	_Ala_	Ser 20	Ile	Ser	Cys	Arg	Ser 25		Gln	Ile	Ile		His O	Ser
45	Asp	Gly	Asn 35	Thr	Tyr	Leu	Glu	Trp	_	Leu	Gln	Lys		Gly 5	Gln	Ser
50	Pro	Lys 50	Leu	Leu	Ile	Tyr	Lys 55		Ser	Asn	Arg	_	Ser 0	Gly	Val	Pro
·	Asp 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Ser 75	Phe	Thr	Leu	Ala	Ile 80

5	Ser Arg Val Glu Ala Glu Asp Glu Gly Val Tyr Tyr Cys Phe Gln Gly 85 90 95
	Ser His Val Pro His Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile 100 105 110
10	(28) INFORMATION FOR SEQ ID NO:28
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 112 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:28:
20	Asp Val Val Met Thr Gln Ser Pro Leu Ser Leu Pro Val Thr Leu Gly 1 5 10 15
25	Gln Pro Ala Ser Ile Ser Cys Arg Ser Ser Gln Ser Leu Val Tyr Ser 20 25 30
·	Asp Gly Asn Thr Tyr Leu Asn Trp Phe Gln Gln Arg Pro Gly Gln Ser 35 40 45
30	Pro Arg Arg Leu Ile Tyr Lys Val Ser Asn Arg Asp Ser Gly Val Pro 50 55 60
35	Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Lys Ile 65 70 75 80
	Ser Arg Val Glu Ala Glu Asp Val Gly Val Tyr Tyr Cys Met Gln Gly 85 90 95
40	Thr His_Trp Ser Trp Thr Phe Gly Gln Gly Thr Lys Val Glu Ile Lys 100 105 110
45	(29) INFORMATION FOR SEQ ID NO:29
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 111 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:29:

5	Asp 1	Val	Leu	Met	Thr 5	Gln	Ser	Pro	Leu	Ser 10	_	Pro	Val	Thr	Leu (_
10	Gln	Pro	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25		Gln	Ile	Ile	Ile :		Ser
•	Asp	Gly	Asn 35	Thr	Tyr	Leu	Glu	Trp		Gln	Gln	Arg	Pro 4	Gly 5	Gln	Ser
15	Pro	Arg 50	Leu	Leu	Ile	Tyr	Lys 55		Ser	Asn	Arg	Phe 6		Gly	Val	Pro
20	Asp 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Asp 75	Phe	Thr	Leu	Lys	Ile 80
	Ser	Arg	Val	Glu	Ala 85		Asp	Val	Gly	Val 9		Tyr	Cys	Phe		Gly 5
25	Ser	His	Val	Pro 100		Thr	Phe	e Gly	Gly 105		Th:	c Ly:	s Vai	1 Glu 110	ı Ile	.
	(30)	INF	ORMA	TION	FOR	SEÇ	ID	NO:	30							
30			(i)	(A) I B) T	ENGT	H: :	TERIS 112 a ino a : lin		S: pac	ids	-			·	
		(ii)	MOLE	CULE	TYP	e: 1	pept	ide							
35		(xi)	SEQU	JENCI	E DES	CRI	PTIO	N: S	EQ I	ои о	:30:				
	Asp 1		Val	. Met		Gln 5	Ser	Pro	Asp	Ser 1	Leu 10	Ala	. Val	Ser	Leu	Gly 15
4 0	Glu	Arg	_Ala	Thr 20		Asn	Cys	Lys	Ser 2	Ser 5	Gln	Ser	· Val	Leu	Tyr 30	Ser
45 ⁻	Ser	: Asr	Asr 35		. Asr	Tyr	: Lev	ı Ala	Trp	туг	Glr	Glr	lys	Pro 45	Gly	Gln
50	Pro	Pro 50		s Lev	ı Lev	ı Ile	Tyn	r Trp 55) Ala	. Šer	Thr	. Arq	y Glu 60	ı Ser	Gly	Val
- -	Pro 65		o Arq	g Phe	e Sei	r Gly	y Sei	r Gl	y Sei	c Gly	7 Thi 7!	: Asj	p Phe	e Thr	Leu	Thr 80

5	Ile	Ser	Ser	Leu	Gln 85	Ala	Glu	Asp	Val	Ala 90		Tyr	Tyr	Cys	_	G1n 95
10	Tyr	Asp	Thr	Ile 100	Pro	Thr	Phe	Gly	Gly 105	Gly 5	Thr	Lys	Val	Glu 11		Lys
	(31)	INF	ORMA	TION	FOR	SEQ	ID	NO: 3	1							
15			(i)	(1	A) L	ENGT YPE:	H: 1 ami	11 a no a	mino cid	: açi	.ds					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
20		(:	ĸi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	31:			•	
	Asp 1	Val	Leu	Met	Thr 5	Gln	Thr	Pro	Asp	Ser 1	_	Pro	Val	Ser	_	Gly .5
25	Asp	Arg	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25		Gln	Ile	Ile		His O	Ser
30	Asp	Gly	Asn 35	Thr	Tyr	Leu	Glu	Trp		Leu	Gln	Lys		Gly 5	Gln	Ser
	Pro	Lys 50	Leu	Leu	Ile	Tyr	Lys 55		Ser	Asn	Arg	_	Ser 0	Gly	<u>V</u> al	Pro
35	Asp 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Asp 75	Phe	Thr	Leu	Met	Ile 80
40	Ser	Arg	Val	Glu	Ala 85	Glu	Asp	Leu	Gly	Val 90	_	Tyr	Cys	Phe	_	Gly 5
	Ser	His	Val	Pro 100	His	Thr	Phe	Gly	Gly 105		Thi	. Ly:	s Le	u Gl		e
45	(32)	INF	ORMA	TION	FOR	SEQ	ID	ио: 3	2							
5 0	r		(i)	(1		ENGT YPE:	H: 1 ami	17 a	mino	: aci	.ds					
50 .		(:	ii)	MOLE	CULE	TYP	E: p	epti	.de							
		(:	xi)	SEQU	ence	DES	CRIP	TION	: SE	Q II	NO:	32:				

5	Asp 1	Val	Gln	Leu	Val 5	Glu	Ser	Gly	Gly	Gly , 1		Val	Gln	Pro		Gly .5
10	Ser	Arg	Lys	Leu 20	Ser	Cys	Ala	Ala	Ser 25		Phe	Thr	Phe	_	Ser 0	Phe
	Gly	Met	His 35	Tṛp	Val	Arg	Gln	Ala 40		Glu	Lys	Gly	_	Glu 5	Trp	Val
15	Ala	Tyr 50		Ser	Ser	Gly	Ser 55		Thr	Ile	Tyr	_	Ala O	Asp	Thr	Val
20	Lys 65	Gly	Arg	Phe	Thr.	Ile 70	Ser	Arg	Asp	Asn	Pro 75	Lys	Asn	Thr	Leu	Phe 80
	Leu	Gln	Met	Thr	Ser 85	Leu	Arg	Ser	Glu	Asp 90		Ala	Met	Tyr		Cys 5
25	Ala	Arg	Met	Arg 100	Lys	Gly	Tyr	Ala	Met 105	_	Tyr	Trp	Gly	Gln 11	_	Thr
30	Thr	Val	Thr 115	Val	Ser											
	(33)	INF	ORMA!	TION	FOR	SEQ	ID	мо: 3	3	-						
35		•	(i) :	(1	A) L: B) T	ENGT YPE:		25 a no a	mino cid		ds					
35		•		(1	A) L: B) T' C) To	ENGT YPE: OPOL	H: 1 ami OGY:	25 a no a lin	mino cid ear		ds					
35 40		(: (:	ii) *i)	() () MOLE SEQUI	A) L: B) T' C) T(CULE ENCE	ENGT YPE: OPOL TYP DES	H: 1 ami OGY: E: p CRIP	25 a no a lin epti TION	mino cid ear de	aci) NO:					
	Glu 1	(: (:	ii) *i)	() () () MOLE	A) L: B) T' C) T(CULE ENCE	ENGT YPE: OPOL TYP DES	H: 1 ami OGY: E: p CRIP	25 a no a lin epti TION	mino cid ear de : SE	aci	NO:		Gln	Pro	Gly 1	Arg .5
	1	(: Val	ii) xi) _Gln_	() () MOLE SEQUI	A) L: B) T C) T CULE ENCE Val	ENGT YPE: OPOL TYP DES Gln	H: 1 ami OGY: E: p CRIP	25 a no a lin epti TION	mino cid ear de : SE	Gly Gly	NO: Val	Val		Ser	_1	.5
40	1 Ser	(; Val Leu	ii) xi) _Gln_ Arg	(1) (1) (1) (1) MOLE SEQUI Leu	A) L: B) T' C) T(CULE ENCE Val 5	ENGT YPE: OPOL TYP DES Gln	H: 1 ami OGY: E: p CRIP Ser	25 a no a lin epti TION Gly	mino cid ear de : SE Gly Ser 25	Gly Gly	Val O Phe	Val	Phe Leu	Ser 3	Ser 0	.5 Tyr

5	Lys Gly Arg Phe Thr Ile Ser Arg Asn Asp Ser Lys Asn Thr Leu Phe 65 70 75 80
	Leu Gln Met Asp Ser Leu Arg Pro Glu Asp Thr Gly Val Tyr Phe Cys 85 90 95
10	Ala Arg Asp Gly Gly His Gly Phe Cys Ser Ser Ala Ser Cys Phe Gly 100 105 110
15	Pro Asp Tyr Trp Gly Gln Gly Thr Pro Val Thr Val Ser 115 120 125
	(34) INFORMATION FOR SEQ ID NO:34
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 117 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:
	Glu Val Gln Leu Val Glu Ser Gly Gly Gly Val Val Gln Pro Gly Arg 1 5 10 15
30	Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Ile Phe Ser Ser Phe 20 25 30
35	Gly Met His Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val 35 40 45
	Ala Tyr Ile Ser Ser Asp Gly Phe Thr Ile Tyr His Ala Asp Ser Val 50 55 60
40	Lys Gly Arg Phe Thr Ile Ser Arg Asp Asp Pro Lys Asn Thr Leu Phe 65 70 75 80
45	Leu Gln Met Thr Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr Cys 85 90 95
	Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr Trp Gly Gln Gly Thr
50	Thr Val Thr Val Ser
	(35) INFORMATION FOR SEQ ID NO:35
55	

5		i	(i) :	(I	ENCE A) Li B) TY C) TO	ENGTI (PE:	H: 1 ami	20 a	mino cid		ds					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de	-						
10		(2	ĸi)	SEQUI	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	35:				
	Gln 1		Gln	Leu	Val 5	Glu	Ser	Gly	Gly	Gly 10	_	Val	Gln	Pro	_	Arg 5
15	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 25	_	Phe	Thr	Phe	Ser 3	_	Tyr
20	Ala	Met	His 35	_	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Val
25	Ala	Val 50	Ile	Ser	Tyr	Asp	Gly 55	_	Asn	Lys	Tyr	_	Ala O	Asp	Ser	Val
	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ser 75	Lys	Asn	Thr	Leu	Tyr 80
30	Leu	Gln	Met	Asn	Ser 85	Leu	Arg	Ala	Glu	Asp 9	_	Ala	Val	Tyr		Cys 5
35	Ala	Arg	Asp	Arg 100		Asp	Trp	Gly	Trp 10		Leu	Phe	Asp	Tyr 11		Gly
	Gln	Gly	Thr 115	Leu	Val	Thr	Va]	Ser 120								
40	(36)	INF	ORMA	TION	FOR	SEQ	ID	NO:3	36		-					
			(i)	(ENCE A) L B) T C) T	ENGT YPE:	H: 1 ami	l17 a ino a	amino acid	ac:	ids				-	
4 5	••		•	MOLE						EO I	D NO	:36:				
50	Gln 1	Val	-		Val					Gly			Gln	Pro	Gly	Arg 15

5	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 25		Phe	Thr	Phe	_	Ser 0	Phe
10	Gly	Met	His 35	Trp	Val	Arg	Gln	Ala 40	Pro	Gly	Lys	Gly		Glu 5	Trp	Val
	Ala	Tyr 50	Ile	Ser	Ser	Gly	Ser 55	Phe	Thr	Ile	Tyr		Ala O	Asp	Ser	Val
15	Lys 65	Gly	Arg	Phe	Ţhr	Ile 70	Ser	Arg	Asp	Asn	Ser 75	Lys	Asn	Thr	Leu	Tyr 80
20	Leu	Gln	Met	Asn	Ser 85	Leu	Arg	Ala	Glu	Asp 9		Ala	Val	Tyr		Cys 5
20	Ala	Arg		Arg 100	Lys	Gly	Tyr	Ala	Met 109		Tyr	Trp	Gly	Gln 11	_	Thr
25	Leu	Val	Thr 115	Val	Ser											
	(37)	INFO	ORMA!	TION	FOR	SEQ	ID	NO: 3	7							٠
30			(i) {	(1	ENCE A) Li B) T'	ENGT:	H: 9 ami	8 am no a	ino cid		ls					•
		. (:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
35		(2	ki) S	SEQUI	ENCE	DES	CRIP	TION	: SE	Q II	NO:	37:				
40	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1		Val	Gln	Pro		Gly .5
	Ser	Leu_	Arq	Leu 20	Ser	Cys	Ala	Ala	Ser 25		Phe	Thr	Phe		Ser 0	Tyr
45	Trp	Met	Ser 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Val
50	Ala	Asn 50	Ile	Lys	Gln	Asp	Gly 55		Glu	Lys	Tyr	_	Val 0	Asp	Ser	Val
	Tura	Gly	3	Dha	mb	Tla	C^=	3	3	3	210	T	۸cn	C	T	Петъ

5	Leu	Gln	Met	Asn	Ser 85	Leu	Arg	Ala	Glu	Asp 90		Ala	Val	Tyr	Tyr 9	Cys 5
•	Ala	Arg														
10	(38)	INF	ORMA'	TION	FOR	SEQ	ID	NO:3	8							
			(i)	(1	A) L: B) T:	ENGT YPE:	H: 1 ami		mino cid	aci	ds					
15		(:	ii)	MOLE	CULE	TYP	E: p	epti	de			٠				
		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SI	EQ. ID	NO:	38:				
20	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1		Val	Gln	Pro		Gly .5
25	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 2		Phe	Thr	Phe		Ser 30	Phe
	Gly	Met	His 35	Trp	Val	Arg	Gln	Ala 4		Gly	Lys	Gly		Glu 15	Trp	Val
30	Ala	Tyr 50		Ser	Ser	Gly	Ser 5		Thr	Ile	Tyr		Ala 50	Asp	Ser	Val
35	Lys 65	_	Arg	Phe	Thr	Ile 70		Arg	Asp	Asn	Ala 75		Asn	Thr	Leu	Phe 80
	Leu	Gln	Met	Thr	Ser 85		Arg	Ala	Glu		Thr O	Ala	Met	Tyr	Tyr	Cys 95
40	Ala	Arg	_Met	Arg 100		Gly	Tyr	Ala	Met 10	: Asp 05	Tyr	Trp	Gly	Gln 1	Gly 10	Thr
45	Thr	' Val	Th:	r Val	Ser	•									•	
	(39)	INE	FORM	OITA	FOF	SE	Q ID	NO:	39							
50			(i)		(A) 1 (B) 7	LENG TYPE	TH: : am	TERI 15 a ino : li	mino acid	aci I	ds					
•			(ii)	MOL	ECUL	E TY	PE:	pept	ide							

		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:39:
5	Met 1	Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr 5 10 15
	(40)	INFORMATION FOR SEQ ID NO:40
10		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:
20	Lys 1	Thr Ser Leu Arg Pro Gly Lys Gly Ser Ser Asp Tyr Glu Lys Lys 5 10 15
	(41)	INFORMATION FOR SEQ ID NO:41
25		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:
	Lys 1	Thr Ser Leu Arg Pro Gly Lys Gly Ser Ser Glu Tyr Glu Lys Lys 5 10 15
35	(42)	INFORMATION FOR SEQ ID NO:42
40		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(Ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:42:
45	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp His Glu Lys Lys 5 10 15
50	(43)	INFORMATION FOR SEQ ID NO:43
		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55		(-/

5	(11) Reddeddd 1111. peperae
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:
10	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Lys Lys 1 5 10 15
	(44) INFORMATION FOR SEQ ID NO:44
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:
	Gln Ser Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Lys Lys 1 5 10 15
25	(45) INFORMATION FOR SEQ ID NO:45
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:45:
35	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glu Lys Lys 1 5 10 15
	(46) INFORMATION FOR SEQ ID NO:46
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:
50	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glx Lys Lys 1 5 10 15
	(47) INFORMATION FOR SEQ ID NO:47
	(i) SEQUENCE CHARACTERISTICS:
55	

5		(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:47:
10	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glu Lys Thr 5 10 15
15	(48)	INFORMATION FOR SEQ ID NO:48
		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20		(ii) MOLECULE TYPE: peptide
	٠	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:48:
25	Gln 1	Thr Ser Leu Arg Ala Asp Lys Gly Ser Ser Asp Gln Glu Lys Lys 5 10 15
	(49)	INFORMATION FOR SEQ ID NO:49
30		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:49:
40	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Lys Ser Asp Ser Glu Lys Lys 5 10 15
70	(50)	INFORMATION FOR SEQ ID NO:50
4 5		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:50:
	Gln 1	Thr Ser Leu Arg Pro Ala Arg Gly Ser Ser Asp Gln Glu Lys Lys 5 10 15

5	(51)	INFORMATION FOR SEQ ID NO:51
·		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:51:
15	Gln 1	Thr Ser Leu Lys Pro Gly Arg Gly Ser Ser Asp Pro Glu Lys Lys 5 10 15
	(52)	INFORMATION FOR SEQ ID NO:52
20		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:
	Gln 1	Thr Ser Leu Arg Pro Gly Arg Gly Ser Ser Asp Thr Glu Lys Lys 5 10 15
30	(53)	INFORMATION FOR SEQ ID NO:53
35		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:
40	Ġln 1	Ile_Ser_Leu Arg Pro Gly Lys Gly Ser Ser Asp Ser Glu Lys Lys 5 10 15
45	(54)	INFORMATION FOR SEQ ID NO:54
4 3	·	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:

5	Gln 1	Thr Ser Leu Arg Pro Gly Lys Gly Asp Ser Asp Glu Asp Lys Lys 5 10 15
	(55)	INFORMATION FOR SEQ ID NO:55
10		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
15		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:55:
	Glu 1	Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asp Ala Asp Lys Lys 5 10 15
20	(56)	INFORMATION FOR SEQ ID NO:56
25		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:56:
30	Val 1	Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asp Glu Asp Lys Lys 5 10 15
35	(57)	INFORMATION FOR SEQ ID NO:57
		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:57:
45	Val 1	Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asp Glu Glu Lys Lys 5 10 15
	(58)	INFORMATION FOR SEQ ID NO:58
50		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:58:
	Val Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asx Ala Asx Lys Lys 1 5 10 15
10	(59) INFORMATION FOR SEQ ID NO:59
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:59:
	Val Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asp Glu Asp Asp Glu 1 5 10 15
25	(60) INFORMATION FOR SEQ ID NO:60
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:60:
35	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Thr Thr 1 5 10 15
((61) INFORMATION FOR SEQ ID NO:61
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:
50	Gln Asn Ser Leu Thr Pro Gly Lys Gly Ser Ser Pro Glu Lys Lys 1 5 10 15
	(62) INFORMATION FOR SEQ ID NO:62
	(i) SEQUENCE CHARACTERISTICS:
55	

5		(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:
10	Val 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Asp Lys Lys 5 10 15
15	(63)	INFORMATION FOR SEQ ID NO:63
		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20		(ii) MOLECULE TYPE: peptide
•		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:63:
25	Val 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
•	(64)	INFORMATION FOR SEQ ID NO:64
30		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:64:
40	Val 1	Thr Arg Val Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
•	(65)	INFORMATION FOR SEQ ID NO:65
45		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:65:
	Leu 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Glu Lys Lys 5 10 15
55		

	(66) INFORMATION FOR SEQ ID NO:00
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:66:
15	Val Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Glu Gln Lys 1 5 10 15
	(67) INFORMATION FOR SEQ ID NO:67
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:67:
	Val Thr Lys Val Arg Pro Glu Lys Gly Asp Ser Asp Ala Glu Lys Lys 1 10 15
30	(68) INFORMATION FOR SEQ ID NO:68
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:68: Val Thr Lys Val Arg Pro Glu Lys Gly Asp Ser Asp Ser Glu Lys Lys
	val Thr Lys val Alg Flo Gla Lys of hop to the 15
45	(69) INFORMATION FOR SEQ ID NO:69
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:69:

5	Val 1	Thr Lys Val Ser Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(70)	INFORMATION FOR SEQ ID NO:70
10		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:70:
20	Val	Thr Lys Val Arg Ser Gly Lys Gly Glu Ser Asp Ala Glu Lys Lys 5 10 15
	(71)	INFORMATION FOR SEQ ID NO:71
25		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
-		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:71:
	Val 1	Thr Ser Val Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
35	(72)	INFORMATION FOR SEQ ID NO:72
40		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(<u>ii</u>) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:72:
45	Val	Ser Ser Val Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
50	(73)	INFORMATION FOR SEQ ID NO:73
55		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

		(ii) MOLECULE TYPE: peptide
5		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:73:
	Val 1	Thr Ser Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
10	(74)	INFORMATION FOR SEQ ID NO:74
15		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
20		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:74:
	Val 1	Ser Ser Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
25	(75)	INFORMATION FOR SEQ ID NO:75
		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:75:
35	Val	Thr Ser Ala Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(76)	INFORMATION FOR SEQ ID NO:76
40		 (i) SEQUENCE CHARACTERISTICS: - (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45		(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:76:
50	Val 1	Ser Pro Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
-	(77)	INFORMATION FOR SEQ ID NO:77
		(i) SEQUENCE CHARACTERISTICS:

5		(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:77:
	Val 1	Thr Lys Ala Arg Pro Gly Lys Gly Asp Ser Asp Val Glu Lys Asn 5 10 15
15	(78)]	INFORMATION FOR SEQ ID NO:78
20		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:78:
25	Val T 1	Thr Leu Ile Pro Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(79) 1	NFORMATION FOR SEQ ID NO:79
30		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:79:
40	Val T	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(80) 1	ENFORMATION FOR SEQ ID NO:80
45		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:80:
	Val 1	Chr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Asp Lys Lys 5 10 15

5	(81)	INFORMATION FOR SEQ ID NO:81
v		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:81:
15	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Arg Lys 5 10 15
	(82)	INFORMATION FOR SEQ ID NO:82
20	,	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	,	(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:82:
	Val 1	Thr Leu Leu Gln Ala Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
30	(83)	INFORMATION FOR SEQ ID NO:83
35		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:83:
40	Val 1	Thr_Leu Leu Gln Pro Gly Glu Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
45	(84)	INFORMATION FOR SEQ ID NO:84
		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:84:

5	Leu 1	Thr Leu Leu Gln Pro Gly Asn Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(85)	INFORMATION FOR SEQ ID NO:85
10		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:85:
20	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Ile 5 10 15
	(86)	INFORMATION FOR SEQ ID NO:86
25		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
•		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:86:
	Val 1	Thr Leu Phe Gln Pro Gly Gln Gly Asp Ser Asp Pro Glu Lys Lys 5 10 15
35	(87)	INFORMATION FOR SEQ ID NO:87
40		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(<u>i</u> i) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:87:
45	Val 1	Thr Leu Pro Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
50	(88)	INFORMATION FOR SEQ ID NO:88
55		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
55		

5		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:88:
10	Val 1	Thr Leu Pro Gln Pro Gly Lys Gly Asp Trp Asp Ala Glu Lys Lys 5 10 15
	(89)	INFORMATION FOR SEQ ID NO:89
15		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	•	(ii) MOLECULE TYPE: peptide
20		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:89:
	Val 1	Thr Phe Leu Ser Pro Gly Gln Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
25	(90)	INFORMATION FOR SEQ ID NO:90
30		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:90:
35	Glu 1	Ser Ser Ala Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(91)	INFORMATION FOR SEQ ID NO:91
40	٠	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:91:
50	Val 1	Thr Leu Ser Ser Pro Gly Gln Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(92)	INFORMATION FOR SEQ ID NO:92
5 5		(i) SEQUENCE CHARACTERISTICS:

5		(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:92:
	Val 1	Thr Thr Ala Lys Pro Glu Lys Gly Asp Ser Asp Val Glu Lys Lys 5 10 15
15	(93)	INFORMATION FOR SEQ ID NO:93
20	·	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:93:
25	Val 1	Thr Thr Pro Lys Pro Asp Lys Gly Asp Ser Asp Val Glu Lys Lys 5 10 15
	(94)	INFORMATION FOR SEQ ID NO:94
30		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:94:
40	Val 1	Thr Ala Pro Arg Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
	(95)	INFORMATION FOR SEQ ID NO:95
45		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:95:
•	Val	Thr Ala Pro Lys Pro Gly Lys Gly Thr Ser Ser Ala Glu Lys Lys 5 10 15

	(96)	INFORMATION FOR SEQ ID NO:96
5		(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:96:
15	Val 1	Thr Thr Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
	(97)	INFORMATION FOR SEQ ID NO:97
20		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:97:
•	Val 1	Ser Ala Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
30	(98)	INFORMATION FOR SEQ ID NO:98
35		(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:98:
40	Val	Thr_Ala Pro Arg Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
45	(99)	INFORMATION FOR SEQ ID NO:99
50		 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	•	(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:99:
		·

5	1 5 10 15
	(100) INFORMATION FOR SEQ ID NO:100
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
,	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:100:
	Val Thr Ala Pro Lys Pro Asp Lys Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
20	(101) INFORMATION FOR SEQ ID NO:101
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:101:
	Val Thr Ala Pro Lys Ser Glu Lys Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
35	(102) INFORMATION FOR SEQ ID NO:102
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	(<u>ii</u>) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:102:
45	Phe Thr Ala Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
	(103) INFORMATION FOR SEQ ID NO:103
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:103:
	Leu Thr Ala Pro Lys Pro Gly Arg Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
10	(104) INFORMATION FOR SEQ ID NO:104
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
• .	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:104:
20	Val Thr Ala Pro Lys Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Arg 1 5 10 15
25	(105) INFORMATION FOR SEQ ID NO:105
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:105:
35	Val Ser Ala Pro Lys Pro Gly Lys Glu Gly Ser Ser Ala Glu Lys Lys 1 10 15
-	(106) INFORMATION FOR SEQ ID NO:106
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:106:
50	Val Thr Ala Pro Lys Pro Arg Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
	(107) INFORMATION FOR SEQ ID NO:107
i 5	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:107:
10	Val Thr Phe Leu Ser Pro Gly Gln Gly Asn Ser Asp Ala Glu Leu Pro 1 5 10 15
	(108) INFORMATION FOR SEQ ID NO:108
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:108:
25	Val Thr Phe Leu Ser Pro Gly Gln Gly Asn Ser Asp Glu Asp Leu Pro 1 5 10 15
	(109) INFORMATION FOR SEQ ID NO:109
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:109:
40	Val Thr Leu Ser Ser Pro Gln Arg Gly Asp Ser Asp Ala Glu Lys Lys 1 5 10 15
	(110) INFORMATION FOR SEQ ID NO:110
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:110:
	Val Thr Ala Pro Lys Ser Ser Lys Gly Gly Ser Ser Ala Glu Lys Lys 1 5 10 15
55	

5	(111) INFORMATION FOR SEQ ID NO:111
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:111:
15	Gln Thr Ser Pro Thr Pro Gly Lys Gly Ser Ser Asp Pro Glu Lys Lys 1 5 10 15
	(112) INFORMATION FOR SEQ ID NO:112
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:112:
30	Gln Ile Ser Leu Ile Pro Gly Lys Gly Ser Tyr Asp Asp Glu Lys Lys 1 5 10 15
	(113) INFORMATION FOR SEQ ID NO:113
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:113:
	Val Thr_Ala Leu Lys Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
45	(114) INFORMATION FOR SEQ ID NO:114
,50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:114:

5	1 5 Lys Gly Ala Ser Ser Gly Glu Lys Lys 1 5 10 15
	(115) INFORMATION FOR SEQ ID NO:115
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:115:
	Val Thr Pro Pro Ser Pro Gly Gln Gly Asp Ser Ala Ala Glu Lys Lys 1 5 10 15
20	(116) INFORMATION FOR SEQ ID NO:116
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:116:
	Val Thr Pro Pro Ser Pro Gly Gln Gly Asp Ser Ala Arg Glu Lys Lys 1 5 10 15
35	(117) INFORMATION FOR SEQ ID NO:117
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid
40	(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:117:
45	Val Thr Val Arg Lys Pro Gly Lys Gly Asp Ser Ser Asp Glu Lys Lys 1 5 10 15
	(118) INFORMATION FOR SEQ ID NO:118
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
5 5	

	(11) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:118:
	Gln Thr Ser Val Arg Leu Gly Gln Gly Ser Ser Asp Pro Glu Lys Lys 1 5 10 15
10	(119) INFORMATION FOR SEQ ID NO:119
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:119:
20	Lys Thr Ser Leu Arg Pro Trp Lys Gly Ser Ser Asp Ser Asp Lys Lys 1 10 15
25	(120) INFORMATION FOR SEQ ID NO:120
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:120:
35	Gln Thr Asp Val Thr Gln Gly Gln Gly Ser Ser Gln Pro Glu Lys Lys 1 5 10 15
	(121) INFORMATION FOR SEQ ID NO:121
40	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:121:
50	Gln Thr Ala Val Ser Gln Gly Gln Gly Ser Ser Gln Ser Glu Lys Lys 1 5 10 15
	(122) INFORMATION FOR SEQ ID NO:122
55	(i) SEQUENCE CHARACTERISTICS:

5	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:122:
10	Leu Thr Ala Pro Arg Thr Asn Arg Gly Ser Ser Asp Ser Glu Lys Lys 1 5 10 15
15	(123) INFORMATION FOR SEQ ID NO:123
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:123:
25	Val Thr Ala Pro Ser Ser His Arg Gly Ser Ser Asp Thr Glu Lys Lys 1 5 10 15
	(124) INFORMATION FOR SEQ ID NO:124
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:124:
40	Leu Leu Ser Leu Ser Pro Leu Lys Gly Asp Ser Asp Pro Glu Lys Val 1 5 10 15
٠	(125) INFORMATION FOR SEQ ID NO:125
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:125:
	Val Thr Ala Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu 1 5 10 15
55	·

	(120) INFORMATION FOR SEQ ID NO:126
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:126:
15	Val Thr Ile Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Let 1 5 10 15
	(127) INFORMATION FOR SEQ ID NO:127
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:127:
	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu 1 5 10 15
30	(128) INFORMATION FOR SEQ ID NO:128
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:128:
	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu 1 10 15
45	(129) INFORMATION FOR SEQ ID NO:129
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:129:
55	

5	Ala Val Ser Pro Thr Pro Asp Thr Gly Val Ile Lys Thr Glu Lys Leu 1 5 . 10 15
	(130) INFORMATION FOR SEQ ID NO:130
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:130:
20	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Pro Ser 1 5 10 15
	(131) INFORMATION FOR SEQ ID NO:131
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:131:
	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Met Lys Leu 1 5 10 15
35	(132) INFORMATION FOR SEQ ID NO:132
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
-	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:132:
45	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Met Arg Leu 1 5 10 15
50	(133) INFORMATION FOR SEQ ID NO:133
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:133:
	Tyr Leu Pro Pro Thr Pro Gly Leu Ile Arg Ser Thr Ser Met Lys Leu 1 5 10 15
10	(134) INFORMATION FOR SEQ ID NO:134
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:134:
	Tyr Leu Pro Pro Thr Pro Gly Leu Ile Arg Ser Thr Ser Val Lys Leu 1 5 10 15
25	(135) INFORMATION FOR SEQ ID NO:135
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
,	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:135:
35	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(136) INFORMATION FOR SEQ ID NO:136
40	 (i) SEQUENCE CHARACTERISTICS: - (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:136:
50	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Gly Lys Leu 1 5 10 15
	(137) INFORMATION FOR SEQ ID NO:137
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGIN: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:137:
10	Tyr Leu Pro Ala Thr Pro Gly Val Val Arg Ser Ser Ala Gly Met Leu 1 5 10 15
15	(138) INFORMATION FOR SEQ ID NO:138
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:138:
25	Ser Leu Pro Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(139) INFORMATION FOR SEQ ID NO:139
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:139:
40	Ser Leu Pro Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Asn Lys Leu 1 5 10 15
	(140) INFORMATION FOR SEQ ID NO:140
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:140:
	Ser Leu Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Glu Lys Leu 1 5 10 15
. 55	

_	(141) INFORMATION FOR SEQ ID NO:141
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:141:
15	Ser Leu Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Asp Lys Leu 1 5 10 15
	(142) INFORMATION FOR SEQ ID NO:142
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:142:
30	Ser Leu Pro Pro Arg Pro Gly Arg Val Arg Ser Ser Ser Glu Lys Leu 1 5 10 15
	(143) INFORMATION FOR SEQ ID NO:143
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:143:
	Ser Leu_Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Glu Gln Leu 1 5 10 15
45	(144) INFORMATION FOR SEQ ID NO:144
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:144:

5	1 5 10 15 15 15 15 15 15 15 15 15 15 15 15 15
	(145) INFORMATION FOR SEQ ID NO:145
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:145:
	Ser Leu Pro Pro Lys Pro Gly Lys Ile Arg Ser Ser Thr Gly Lys Leu 1 5 10 15
20	(146) INFORMATION FOR SEQ ID NO:146
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:146:
	Ser Leu Pro Pro Lys Pro Gly Arg Ile Arg Ser Ser Thr Gly Lys Leu 1 10 15
35	(147) INFORMATION FOR SEQ ID NO:147
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:147:
45	Ser Leu Pro Pro Lys Pro Gly Lys Ile Arg Ser Ser Thr Gly Gln Leu 1 5 10 15
	(148) INFORMATION FOR SEQ ID NO:148
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:148:
	Ser Leu Pro Pro Glu Pro Gly Lys Ile Arg Ser Ser Thr Gly Arg Leu 1 5 10 15
10	(149) INFORMATION FOR SEQ ID NO:149
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:149:
20	Ser Leu Ala Pro Ser Pro Gly Lys Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
25	(150) INFORMATION FOR SEQ ID NO:150
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:150:
35	Ser Leu Pro Pro Arg Pro Gly Lys Ile Arg Ser Ser Thr Gly Asn Val 1 5 10 15
	(151) INFORMATION FOR SEQ ID NO:151
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:151:
50	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(152) INFORMATION FOR SEQ ID NO:152
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:152:
	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Asp Lys Leu 1 5 10 15
15	(153) INFORMATION FOR SEQ ID NO:153
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:153:
25	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Asn Leu 1 5 10 15
	(154) INFORMATION FOR SEQ ID NO:154
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:154:
40	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Ala Val Glu Lys Leu 1 5 10 15
	(155) INFORMATION FOR SEQ ID NO:155
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 15 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50 `	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:155:
	Ser Leu Pro Pro Arg Pro Gly Lys Arg Ser Ser Ala Glu Lys Leu 1 5 10 15

	(156) INFORMATION FOR SEQ ID NO:156
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:156:
15	Ser Leu Ala Pro Ser Pro Gly Lys Val Arg Ser Thr Val Glu Arg Leu 1 5 10 15
	(157) INFORMATION FOR SEQ ID NO:157
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:157:
	Ser Leu Ala Pro Ser Pro Asp Lys Ile Arg Ser Thr Pro Asp Lys Leu 1 5 10 15
30	(158) INFORMATION FOR SEQ ID NO:158
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:
	Ser Leu_Ala Leu Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
45	(159) INFORMATION FOR SEQ ID NO:159
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
-	
00	(ii) MOLECULE TYPE: peptide
55	(ii) MOLECULE TYPE: peptide(xi) SEQUENCE DESCRIPTION: SEQ ID NO:159:

5	1 5 10 15
5	(160) INFORMATION FOR SEQ ID NO:160
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:160:
	Ser Leu Ala Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Tyr Leu 1 5 10 15
20	(161) INFORMATION FOR SEQ ID NO:161
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:161:
30	Ser Leu Pro Leu Thr Pro Gly Leu Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
35	(162) INFORMATION FOR SEQ ID NO:162
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:162:
45	Ser Leu Pro Leu Thr Pro Arg Val Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(163) INFORMATION FOR SEQ ID NO:163
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:163:
	Phe Leu His Pro Thr Pro Gly Thr Asp Ser Ser Ser Thr Glu Lys Leu 1 5 10 15
10	(164) INFORMATION FOR SEQ ID NO:164
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:164:
	Phe Leu Leu Pro Thr Pro Gly Thr Asp Ser Ser Ser Thr Glu Arg Leu 1 5 10 15
25	(165) INFORMATION FOR SEQ ID NO:165
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
00	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:165:
35	Phe Leu His Pro Thr Arg Val Thr Asp Ser Ser Ser Thr Glu Lys Leu 1 5 10 15
	(166) INFORMATION FOR SEQ ID NO:166
40	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:166:
50	Leu Leu Pro Pro Thr Pro Gly Thr Asn Ser Ser Asn Asp Lys Leu 1 5 10 15
	(167) INFORMATION FOR SEQ ID NO:167
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:167:
10	Val Leu Pro Leu Ser Pro His Arg Ile Arg Ser Glu Ser Glu Asn Leu 1 5 10 15
15	(168) INFORMATION FOR SEQ ID NO:168
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:168:
25	Ser Leu Ala Pro Ser Pro Ala Lys Phe Arg Ser Thr Ala Glu Arg Asp 1 5 10 15
	(169) INFORMATION FOR SEQ ID NO:169
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:169:
	Val Thr Ala Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
40	1 20 10
	(170) INFORMATION FOR SEQ ID NO:170
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:170:
	Val Thr Ala Pro Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
55	

	(171) INFORMATION FOR SEQ ID NO.171
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:171:
15	Val Thr Gly Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(172) INFORMATION FOR SEQ ID NO:172
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:172:
	Val Thr Gly Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Asp Lys Lys 1 5 10 15
30	(173) INFORMATION FOR SEQ ID NO:173
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:173:
	Val Thr Gly Pro Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
45	(174) INFORMATION FOR SEQ ID NO:174
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
٠	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:174:

5	1 5 10 15
	(175) INFORMATION FOR SEQ ID NO:175
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:175:
	Val Thr Ala Pro Arg Pro Gly Arg Ile Arg Ser Glu Ser Glu Arg Lys 1 5 10 15
20	(176) INFORMATION FOR SEQ ID NO:176
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
00	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:176:
30	Val Thr Gly Pro Ser Arg Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
35	(177) INFORMATION FOR SEQ ID NO:177
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
-	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:177:
45	Val Thr Val Pro Arg Pro Ser Arg Ile Arg Ser Glu Ser Glu Arg Lys 1 5 10 15
	(178) INFORMATION FOR SEQ ID NO:178
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:178:
	Val Thr Ala Pro Gly Pro Gly Arg Ile Arg Ser Glu Ser Glu Arg Lys 1 10 15
10	(179) INFORMATION FOR SEQ ID NO:179
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:179:
20	Gln Thr Ser Val Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Arg Lys 1 10 15
25	(180) INFORMATION FOR SEQ ID NO:180
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide •
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:180:
35	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Arg Lys 1 5 10 15
	(181) INFORMATION FOR SEQ ID NO:181
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(Xi) SEQUENCE DESCRIPTION: SEQ ID NO:181:
50	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(182) INFORMATION FOR SEQ ID NO:182
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
•	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:182:
10	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
	(183) INFORMATION FOR SEQ ID NO:183
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:183:
25	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
	(184) INFORMATION FOR SEQ ID NO:184
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:184:
4 0	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ala Glu Pro Glu Lys Lys 1 5 10 15
	(185) INFORMATION FOR SEQ ID NO:185
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:185:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asx Pro Glx Lys Lys 1 10 15

(186) INFORMATION FOR SEQ ID NO:186

5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:186:
15	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Asx Lys Lys 1 5 10 15
	(187) INFORMATION FOR SEQ ID NO:187
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(Xi) SEQUENCE DESCRIPTION: SEQ ID NO:187:
	Gln Thr Ser Val Arg Pro Gly Gln Val Arg Ser Asp Pro Glu Arg Lys 1 10 15
30	(188) INFORMATION FOR SEQ ID NO:188
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:188:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser His Pro Glu Lys Lys 1 5 10 15
4 5	(189) INFORMATION FOR SEQ ID NO:189
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:189:

5	In the Ser Val Arg Pro Gly Asn Val Arg Ser Asp Pro Asp Lys Lys 1 5 10 15
	(190) INFORMATION FOR SEQ ID NO:190
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:190:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Thr 1 5 10 15
20	(191) INFORMATION FOR SEQ ID NO:191
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:191:
30	Gln Thr Ser Val Arg Pro Gly Thr Val Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
35	(192) INFORMATION FOR SEQ ID NO:192
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:192:
4 5	Gln Thr Ser Val Arg Pro Glu Lys Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
	(193) INFORMATION FOR SEQ ID NO:193
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
e e	

	(II) MOLECULE TIPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:193:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Ser Asp Lys Lys 1 5 10 15
10	(194) INFORMATION FOR SEQ ID NO:194
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:194:
20	Gln Thr Ser Val Arg Pro Gly Glu Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
25	(195) INFORMATION FOR SEQ ID NO:195
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:195:
35	Gln Thr Ser Val Arg Pro Gly Asx Val Arg Ser Asx Pro Glx Arg Lys 1 5 10 15
	(196) INFORMATION FOR SEQ ID NO:196
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:196:
50	Gln Thr Ser Val Ser Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(197) INFORMATION FOR SEQ ID NO:197
5 5	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:197:
10	Gln Thr Ser Val Arg Pro Gly Lys Val Asn Ser Asp Pro Glu Lys Lys 1 5 10 15
15	(198) INFORMATION FOR SEQ ID NO:198
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:198:
25	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Asp Thr Lys 1 5 10 15
	(199) INFORMATION FOR SEQ ID NO:199
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:199:
40	Gln Thr Ser Val Arg Pro Lys Lys Val Arg Ser Asp Pro Glx Lys Lys 1 5 10 15
•	(200) INFORMATION FOR SEQ ID NO:200
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:200:
	Gln Thr Ser Val Arg Pro Lys Lys Val Arg Phe Asp Pro Glu Lys Lys 1 5 10 15

	(201) INFORMATION FOR SEQ ID NO:201
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:201:
15	Gln Thr Ser Val Arg Ser Gly Lys Val Arg Ser Glu Pro Glu Thr Lys 1 10 15
	(202) INFORMATION FOR SEQ ID NO:202
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:202:
	Val Thr Asn Leu Arg Pro Gly Lys Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
30	(203) INFORMATION FOR SEQ ID NO:203
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:203:
	Val Thr Asp Leu Arg Pro Gly Lys Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
45	(204) INFORMATION FOR SEQ ID NO:204
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 204:

5	1 5 10 15
	(205) INFORMATION FOR SEQ ID NO:205
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:205:
	Lys Thr Ser Val Thr Pro Gly Lys Phe Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
20	(206) INFORMATION FOR SEQ ID NO:206
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:206: Val Thr Leu Leu Pro Pro Gly Arg Val Arg Ser Asp Ala Glu Lys Lys 1 10 15
	1 5 10 13
35	(207) INFORMATION FOR SEQ ID NO:207
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	` '
	(ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:207:
45	Val Thr Leu Leu Pro Pro Gly Glu Val Arg Ser Asp Ala Glu Lys Lys
40	1 5 10 15
	(208) INFORMATION FOR SEQ ID NO:208
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
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	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:208:
	Val Thr Leu Pro Pro Pro Gly Glx Val Arg Ser Asp Ala Glu Arg Lys 1 5 10 15
10	(209) INFORMATION FOR SEQ ID NO:209
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:209:
20	Val Thr Leu Pro Pro Pro Gly Glx Val Arg Ser Asx Ala Glx Asn Lys 1 5 10 15
25	(210) INFORMATION FOR SEQ ID NO:210
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:210:
35	Val Thr Leu Pro Pro Pro Gln Gln Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
	(211) INFORMATION FOR SEQ ID NO:211
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:211:
50	Val Thr Leu Pro Pro Pro Gly Gln Val Thr Ser Asp Ala Glu Lys Lys 1 5 10 15
	(212) INFORMATION FOR SEQ ID NO:212
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:212:
10	Val Thr Leu Pro Pro Ala Gly Gln Val Arg Ser Asp Ala Glu Lys Arg 1 5 10 15
15	(213) INFORMATION FOR SEQ ID NO:213
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 16 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:213:
25	Ala Leu Ser Pro Ser Ser Gly Gln Ser Ser Ser Ala Ser Glu Arg Leu 1 5 10 15
	(214) INFORMATION FOR SEQ ID NO:214
·30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:214:
40	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
	Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 25
45	(215) INFORMATION FOR SEQ ID NO:215
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:215:
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Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val 5 Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 (216) INFORMATION FOR SEQ ID NO:216 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:216: Glu Lys Val Gly Gly Leu Gln Pro Gly Thr Gly Ala Pro Gly Lys Ala 20 Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 25 (217) INFORMATION FOR SEQ ID NO:217 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:217: 35 Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 40 20 (218) INFORMATION FOR SEQ ID NO:218 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids(B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:218:

Glu Lys Met Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 5 Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser (219) INFORMATION FOR SEQ ID NO:219 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids(B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:219: Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 15 Ser Lys Gly Thr Ser Gln Arg Ala Glu Ser 25 (220) INFORMATION FOR SEQ ID NO:220 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:220: 35 Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 40 20 (221) INFORMATION FOR SEQ ID NO:221 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:221:

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5	1 5 10 15
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr
10	(222) INFORMATION FOR SEQ ID NO:222
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:222:
20	Glu Asn Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 20 25
	(223) INFORMATION FOR SEQ ID NO:223
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:223:
	Glu Lys Val Gly Gly Leu Gln Ser Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
40	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr
	(224) INFORMATION FOR SEQ ID NO:224
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:224:

5	1 5 10 15 15 15 16 16 16 16 16 16 16 16 16 16 16 16 16
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Ser 20 25
10 (225) INFORMATION FOR SEQ ID NO:225
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
,,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:225:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Ile Ser Gln Arg Ala Glu Arg 20 25
(226) INFORMATION FOR SEQ ID NO:226
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:226:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ser 1 5 10 15
40	Ala Lys Gly Asx Ser Glx Arg Ala Gln Ser
(227) INFORMATION FOR SEQ ID NO:227
45 [*]	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:227:

Glu Lys Val Gly Gly Leu Gln Pro Gly Ser Gly Thr Pro Gly Lys Ala 5 Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 10 (228) INFORMATION FOR SEQ ID NO:228 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 15 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:228: 20 Glu Lys Val Gly Gly Leu Gln Pro Gly Ser Gly Thr Pro Gly Lys Ala Ser Lys Gly Ser Ser Gln Arg Ala Glu Ser 25 (229) INFORMATION FOR SEQ ID NO:229 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:229: Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Arg Lys Ala 40 Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 20 (230) INFORMATION FOR SEQ ID NO:230 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 50 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:230:

. 5	1 5 10 15
	Ser Lys Gly Asn Ser Gln Arg Pro Asp Ser 20 25
10	(231) INFORMATION FOR SEQ ID NO:231
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:231:
20	Glu Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
	(232) INFORMATION FOR SEQ ID NO:232
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:232:
•	Glu Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr
	(233) INFORMATION FOR SEQ ID NO:233
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:233:

5	1 5 10 15 15 15 16 17 17 17 17 17 17 17 17 17 17 17 17 17
	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
10	(234) INFORMATION FOR SEQ ID NO:234
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:234:
20	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Lys Arg Ser Glu Thr 20 25
	(235) INFORMATION FOR SEQ ID NO:235
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:235:
	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(236) INFORMATION FOR SEQ ID NO:236
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:236:

5	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Asp Lys Asp 1 5 10 15
	Asn Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(237) INFORMATION FOR SEQ ID NO:237
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:237:
20	Glu Lys Val Gly Gly Leu Thr Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
25	Ser Lys Gly Asn Gly Arg Arg Ser Glu Thr 20 25
	(238) INFORMATION FOR SEQ ID NO:238
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:238:
	Glu Met Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
40	Ser Lys Gly Asn Asp Arg Arg Ser Glu Thr
	(239) INFORMATION FOR SEQ ID NO:239
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:239:

	1 5 10 15 ASP
5	Ser Lys Gly Asn Asp Lys Arg Ser Glu Thr 20 25
	(240) INFORMATION FOR SEQ ID NO:240
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:240:
20	Glu Met Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Lys Arg Ser Glu Thr 20 25
25	(241) INFORMATION FOR SEQ ID NO:241
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:241:
35	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(242) INFORMATION FOR SEQ ID NO:242
4 5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
5 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:242:

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5	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(243) INFORMATION FOR SEQ ID NO:243
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:243:
20	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 10 15
25	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(244) INFORMATION FOR SEQ ID NO:244
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:244:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(245) INFORMATION FOR SEQ ID NO:245
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:245:

5	1 5 10 15
	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr
10	(246) INFORMATION FOR SEQ ID NO:246
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:246:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
20	(247) INFORMATION FOR SEQ ID NO:247
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:247:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Ser Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(248) INFORMATION FOR SEQ ID NO:248
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:248:

Asp Lys Met Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 5 Ser Lys Gly Asn Ala Lys Gln Ser Glu Thr 20 (249) INFORMATION FOR SEQ ID NO:249 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:249: Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Asp Lys Asp 20 Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25 (250) INFORMATION FOR SEQ ID NO:250 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:250: 35 Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 40 Ser Lys Gly Asn Ala Glu Lys Ser Glu Thr 20 (251) INFORMATION FOR SEQ ID NO:251 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:251:

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5	1 5 10 15
	Thr Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
10	(252) INFORMATION FOR SEQ ID NO:252
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:252:
20	Glu Asn Val Gly Asp Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
	(253) INFORMATION FOR SEQ ID NO:253
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:253:
	Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Asp Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(254) INFORMATION FOR SEQ ID NO:254
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:254:

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5	1 5 10 17 Fro Gly Lys Gly Thr Pro Glu Lys Asp
	Ser Lys Gly Asn Ala Lys Lys Ser Gly Thr
10	(255) INFORMATION FOR SEQ ID NO:255
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:255:
20	Asp Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
25	Thr Lys Gly Asn Pro Lys Arg Ser Glu Thr 20 25
	(256) INFORMATION FOR SEQ ID NO:256
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:256:
	Asp Gln Val Gly Gly Leu Gln Pro Gly Gln Gly Thr Pro Glu Lys Asn 1 5 10 15
40	Thr Lys Gly Asn Pro Lys Arg Ser Asp Thr
	(257) INFORMATION FOR SEQ ID NO:257
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:257:
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5	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Glu Lys Asp 1 5 10 15
	Ile Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(258) INFORMATION FOR SEQ ID NO:258
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:258:
20	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Arg Thr Pro Glu Lys Asp 1 5 10 15
25	Asn Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
25	(259) INFORMATION FOR SEQ ID NO:259
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:259:
	Asp Lys Val Gly Gly Leu Lys Leu Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(260) INFORMATION FOR SEQ ID NO:260
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:260:

Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 5 Ser Lys Gly Asn Ala Asn Thr Ser Glu Thr 10 (261) INFORMATION FOR SEQ ID NO:261 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:261: 20 Glu His Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp Ser Lys Gly Asn Ala Gly Arg Ser Glu Thr 25 (262) INFORMATION FOR SEQ ID NO:262 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:262: Glu Gln Val Gly Gly Leu Gln Pro Gly Asn Gly Thr Pro Glu Lys Asp 10 40 Thr Thr Gly Asn Ala Lys Arg Ser Glu Thr 20 (263) INFORMATION FOR SEQ ID NO:263 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:263:

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Glu Lys Glu Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Glu Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr (264) INFORMATION FOR SEQ ID NO:264 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:264: Glu Lys Glu Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Glu 20 Ser Lys Gly Asp Ser Lys Arg Pro Glu Thr 20 25 (265) INFORMATION FOR SEQ ID NO:265 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:265: 35 Glu Lys Glu Gly Gly Leu Gln Pro Gly Lys Gly Ser Pro Glu Lys Glu 40 Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 (266) INFORMATION FOR SEQ ID NO:266 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 266:

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Glu Lys Asp Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 5 Ser Lys Gly Asp Ser Lys Arg Val Glu Met 20 (267) INFORMATION FOR SEQ ID NO:267 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:267: Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Glu Lys Asp 20 Thr Thr Gly Asp Ala Gln Arg Ser Glu Thr 20 25 (268) INFORMATION FOR SEQ ID NO:268 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:268: 35 Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Glu Lys Asp Thr Thr Gly Asn Ala Lys Gly Ser Glu Thr 40 20 (269) INFORMATION FOR SEQ ID NO:269 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:269:

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5	1 5 10 15
	Ser Lys Gly Asn Ala Lys Thr Ser Glu Thr 20 25
10	(270) INFORMATION FOR SEQ ID NO:270
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:270:
20	Ser Asp Gln Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
25	Thr Lys Gly Asn Ala Arg Arg Ser Glu Ser 20 25
	(271) INFORMATION FOR SEQ ID NO:271
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:271:
÷	Glu Lys Ile Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
40	Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr
	(272) INFORMATION FOR SEQ ID NO:272
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:272:

Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 5 Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 (273) INFORMATION FOR SEQ ID NO:273 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:273: Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 20 Phe Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 25 (274) INFORMATION FOR SEQ ID NO:274 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:274: . 35 Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Leu 40 Met Lys Glu Asn Ala Lys Arg Ser Glu Thr 20 (275) INFORMATION FOR SEQ ID NO:275 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:275:

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5	Glu Asn Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Leu 1 5 10 15
	Lys Xaa Glu Asn Ala Lys Arg Pro Glu Thr 20 25
10	(276) INFORMATION FOR SEQ ID NO:276
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:276:
20	Glu Lys Leu Gly Gly Leu Gln Pro Gly Asn Gly Asp Leu Gly Lys Pro 1 5 15
25	Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 25
	(277) INFORMATION FOR SEQ ID NO:277
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:277:
	Glu Lys Leu Gly Pro Leu Gln Leu Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
40	Ser Lys Asp Asp Ala Lys Arg Ser Glu Thr 20 25
	(278) INFORMATION FOR SEQ ID NO:278
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
5 0	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:278:

5	1 5 10 15
	Ser Lys Asp Asn Asp Lys Arg Ser Glu Thr 20 25
10	(279) INFORMATION FOR SEQ ID NO:279
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:279:
20	Glu Gln Leu Gly Gly Leu Gln Pro Gly Gly Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Asp Asp Lys Arg Ser Glu Thr 20 25
	(280) INFORMATION FOR SEQ ID NO:280
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:280:
	Glu Gln Val Gly Gly Leu Lys Ala Arg Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Thr Thr Gly Asn Ala Lys Arg Ser Glu Thr 20 25
	(281) INFORMATION FOR SEQ ID NO:281
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:281:

5	Glu Met Val Gly Val Leu Glu Pro Gly Lys Gly Thr Pro Glu Lys Arg 1 5 10 15
10	Gln Glu Gly Asn Ala Lys Arg Ser Glu Thr 20 25 (282) INFORMATION FOR SEQ ID NO:282
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:282:
20	Glu Gln Val Gly Gly Leu Gln Pro Lys Lys Gly Ser Pro Gly Lys Asp 1 5 10 15
25	Ser Lys Asp Asp Ser Gln Lys Thr Glu Thr
•	(283) INFORMATION FOR SEQ ID NO:283
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:283:
	Glu Gln Val Gly Gly Leu Gln Pro Lys Lys Gly Ser Pro Gly Lys Asp 1 5 10 15
40	Ser Lys Asp Asp Ser Gln Lys Thr Glu Arg 20 25
	(284) INFORMATION FOR SEQ ID NO:284
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:284:

Gln Gln Val Pro Glu Leu Lys Pro Gly Arg Gly Thr Pro Gly Lys Glu 5 Asp Lys Gly Thr Ser Ala Arg Asn Asp Thr 20 (285) INFORMATION FOR SEQ ID NO:285 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:285: Gln Gln Val Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Asp 20 Asp Lys Gly Thr Ser Ala Lys Asn Glu Thr 20 25 (286) INFORMATION FOR SEQ ID NO:286 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 286: 35 Gln Gln Val Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Asp Asp Lys Gly Thr Ser Ala Lys Asn Glu Met _ 20 (287) INFORMATION FOR SEQ ID NO:287 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 45 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:287:

Gln Gln Lys Pro Glu Leu Lys Pro Gly Lys Gly Ser Pro Gly Gln Glu 5 Lys Lys Gly Thr Ser Ser Thr Ser Glu Thr 20 10 (288) INFORMATION FOR SEQ ID NO:288 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:288: 20 Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25 (289) INFORMATION FOR SEQ ID NO:289 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:289: 35 Glu Gln Gln Pro Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 5 10 40 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser _ _ 20 (290) INFORMATION FOR SEQ ID NO:290 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid(C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:290:

5	1 5 10 15
	Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25
10	(291) INFORMATION FOR SEQ ID NO:291
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:291:
20	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Gln 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(292) INFORMATION FOR SEQ ID NO:292
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:292:
	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(293) INFORMATION FOR SEQ ID NO:293
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:293:

Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 5 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 (294) INFORMATION FOR SEQ ID NO:294 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:294: Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 20 15 Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25 (295) INFORMATION FOR SEQ ID NO:295 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:295: 35 Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln Lys Lys Gly Lys Ser Ser Thr Phe Glu Ser 40 (296) INFORMATION FOR SEQ ID NO: 296 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:296:

5	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln 1 5 10 15
10	Lys Gln Gly Lys Ser Ser Thr Phe Glu Ser 20 25
	(297) INFORMATION FOR SEQ ID NO:297
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:297:
20	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Glu 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
	(298) INFORMATION FOR SEQ ID NO:298
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:298:
,	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(299) INFORMATION FOR SEQ ID NO:299
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:299:

Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln 5 Lys Lys Ser Asn Ser Ser Thr Ser Glu Ser (300) INFORMATION FOR SEQ ID NO:300 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:300: Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Glu 20 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser (301) INFORMATION FOR SEQ ID NO:301 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:301: 35 Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Glu 1 10 Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 20 25 (302) INFORMATION FOR SEQ ID NO:302 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:302:

5	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Val Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 20 25
10	(303) INFORMATION FOR SEQ ID NO:303
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:303:
	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(304) INFORMATION FOR SEQ ID NO:304
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:304:
	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly 1 5 10 15
40	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser
	(305) INFORMATION FOR SEQ ID NO:305
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:305:

Glu Gln Gln Pro Glu Ala Lys Pro Gly Lys Gly Thr His Gly Lys Gln 5 Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 20 (306) INFORMATION FOR SEQ ID NO:306 10. (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:306: Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Glu 20 1 10 Lys Lys Asp Lys Ser Ser Thr Ser Asp Ser 20 25 25 (307) INFORMATION FOR SEQ ID NO:307 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:307: 35 Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Gly 1 15 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 (308) INFORMATION FOR SEQ ID NO:308 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:308:

. Gln Gln Gln Ala Glu Leu Lys Pro Gly Arg Gly Thr Pro Gly Gln Glu Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 10 (309) INFORMATION FOR SEQ ID NO:309 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 15 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:309: 20 Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 10 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 25 (310) INFORMATION FOR SEQ ID NO:310 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:310: Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 40 Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 (311) INFORMATION FOR SEQ ID NO:311 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:311:

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Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly His Glu 5 Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 (312) INFORMATION FOR SEQ ID NO:312 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid(C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:312: Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 20 Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25 25 (313) INFORMATION FOR SEQ ID NO:313 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:313: 35 Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly His Glu Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser 40 _ 20 25 (314) INFORMATION FOR SEQ ID NO:314 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:314:

	1 5 10 Thr Pro Gly His Glu
5	
•	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(315) INFORMATION FOR SEQ ID NO:315
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:315:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
25	Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
20	(316) INFORMATION FOR SEQ ID NO:316
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:316:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25
	(317) INFORMATION FOR SEQ ID NO:317
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:317:

5	1 5 10 15
	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(318) INFORMATION FOR SEQ ID NO:318
70	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:318:
20	Glu Gln Gln Val Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
25	(319) INFORMATION FOR SEQ ID NO:319
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:319:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Gln Gly Thr Ser Ser Thr Ser Glu Ser 20 25
	(320) INFORMATION FOR SEQ ID NO:320
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:320:
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Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly His Asp 5 Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 10 (321) INFORMATION FOR SEQ ID NO: 321 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:321: 20 Gln Gln Gln Ala Glu Val Arg Pro Gly Lys Gly Thr Pro Gly His Glu Lys Lys Gly Arg Ser Ser Thr Ser Glu Ser 20 25 . (322) INFORMATION FOR SEQ ID NO:322 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:322: Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 10 40 Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser _ 20 (323) INFORMATION FOR SEQ ID NO:323 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:323:

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5	1 5 10 11 Fro Gly Gln Gln 15
	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
10	(324) INFORMATION FOR SEQ ID NO:324
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:324:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Asp Ser 20 25
	(325) INFORMATION FOR SEQ ID NO:325
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 325:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Ser Pro Gly Gln Gln 1 5 10 15
4 0	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser
	(326) INFORMATION FOR SEQ ID NO:326
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:326:

5	Gln His Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Lys Lys Asn Lys Ser Ser Thr Ser Glu Ser 25
10	(327) INFORMATION FOR SEQ ID NO:327
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:327:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Asn Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25 (328) INFORMATION FOR SEQ ID NO:328
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:328:
	Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Ile Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(329) INFORMATION FOR SEQ ID NO:329
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
5 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:329:

Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 (330) INFORMATION FOR SEQ ID NO:330 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:330: Gln Gln Gln Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 20 Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 25 (331) INFORMATION FOR SEQ ID NO:331 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:331: 35 Gln Gln Gln Thr Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 15 40 Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 (332) INFORMATION FOR SEQ ID NO:332 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:332:

Glu Gln Gln Ala Glu Leu Arg Thr Gly Lys Gly Thr Pro Gly Gln Glu 5 Arg Lys Gly Lys Ser Ser Thr Ser Glu Ser (333) INFORMATION FOR SEQ ID NO:333 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:333: Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 20 Lys Lys Asp Lys Ser Ser Thr Phe Glu Ser 20 25 (334) INFORMATION FOR SEQ ID NO:334 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:334: 35 Glu Gln Gln Ala Glu Leu Arg Pro Gly Thr Gly Ala Pro Gly Gln Glu Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 40 20 (335) INFORMATION FOR SEQ ID NO:335 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 . (xi) SEQUENCE DESCRIPTION: SEQ ID NO:335:

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5	Gln Gln Gln Pro Glu Val Arg Pro Gly Lys Gly Thr His Ala Lys Gln 1 5 10 15
	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(336) INFORMATION FOR SEQ ID NO:336
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:336:
20	Gln Gln Gln Pro Glu Val Arg Pro Gly Lys Asp Thr His Ala Lys Gln 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(337) INFORMATION FOR SEQ ID NO:337
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:337:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Glu Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(338) INFORMATION FOR SEQ ID NO:338
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide(xi) SEQUENCE DESCRIPTION: SEQ ID NO:338:

Glu Gln Gln Thr Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 5 Lys Lys Gly Arg Ser Ser Thr Ser Glu Ala 20 (339) INFORMATION FOR SEQ ID NO:339 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:339: 20 Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 1 Lys Lys Ser Lys Pro Ser Thr Ser Glu Ser 20 25 (340) INFORMATION FOR SEQ ID NO:340 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 35 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:340: Gln Gln Gln Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu Lys Lys Ser Lys Pro Ser Thr Ser Glu Ser 20 (341) INFORMATION FOR SEQ ID NO:341 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:341:

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5	Gin Gin Arg Ala Giu Leu Lys Pro Gly Lys Asp Thr Pro Gly Arg Glu 1 5 10 15
Ū	Lys Lys Asn Lys Pro Ser Thr Ser Glu Ser 20 25
10	(342) INFORMATION FOR SEQ ID NO:342
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:342:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 1 5 10 15
25	Lys Lys Ser Thr Ser Ser Thr Ser Glu Ser 20 25
	(343) INFORMATION FOR SEQ ID NO:343
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:343:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Ser Thr Ser Ser Thr Ser Asp Ser
	(344) INFORMATION FOR SEQ ID NO:344
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:344:

Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Ile Gln Gln 5 Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 (345) INFORMATION FOR SEQ ID NO:345 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:345: Gln Gln Gln Ala Glu Phe Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 20 15 His Arg Ser Lys Pro Ser Thr Ser Glu Ser 20 25 25 (346) INFORMATION FOR SEQ ID NO:346 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:346: 35 Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Leu Gly Gln Glu 1 Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 40 20 (347) INFORMATION FOR SEQ ID NO:347 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:347:

5	Gln Gln Gln Pro Glu Val Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly 1 5 10 15
	Asn Thr Asp Lys Ser Ser Thr Ser Glu Ser 20 25
10	(348) INFORMATION FOR SEQ ID NO:348
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:348:
20	Glu Gln Gln Ala Glu Val Arg Ala Gly Lys Gly Ser Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(349) INFORMATION FOR SEQ ID NO:349
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:349:
	Gln Gln Leu Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
40	Lys Lys Gly Ile Ser Ser Thr Ser Glu Ser 20 25
	(350) INFORMATION FOR SEQ ID NO:350
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:350:

5	1 5 10 15 15
	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(351) INFORMATION FOR SEQ ID NO:351
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:351:
20	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Arg Asn Gly Lys Glu 1 5 10 15
25	Asn Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(352) INFORMATION FOR SEQ ID NO:352
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:352:
	Gln Gln Gln Thr Glu Leu Arg Pro Gly Arg Gly Thr Thr Gly Gln Glu 1 5 10 15
40	Arg Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(353) INFORMATION FOR SEQ ID NO:353
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:353:

Gln His Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu Asn Lys Val Thr Ser Ser Thr Ser Glu Ser (354) INFORMATION FOR SEQ ID NO:354 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
(B) TYPE: amino acid
(C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:354: Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 20 Gln Lys Ala Lys Ser Ser Thr Ser Glu Ser 20 25 (355) INFORMATION FOR SEQ ID NO:355 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:355: 35 Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 Lys Thr Gly Thr Ser Ser Thr Thr Glu Ser 40 20 (356) INFORMATION FOR SEQ ID NO:356 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:356:

5	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Gly Gln Glu 1 5 10 15
	Lys Lys Ser Thr Ser Ser Ala Ser Glu Ser 20 25
10	(357) INFORMATION FOR SEQ ID NO:357
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
•	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:357:
20	Glu Gln Gln Thr Val Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Lys Gly Thr Ser Ala Thr Asn Glu Ser 20 25
	(358) INFORMATION FOR SEQ ID NO:358
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:358:
	Gln Gln Leu Thr Glu Leu Lys Pro Gly Asn Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 25
	(359) INFORMATION FOR SEQ ID NO:359
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:359:

5	1 5 10 15
	Lys Lys Gly Thr Ser Ser Thr Ser Lys Ser 20 25
10	(360) INFORMATION FOR SEQ ID NO:360
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
,,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:360:
20	Leu Gln Gln Pro Val Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
	(361) INFORMATION FOR SEQ ID NO:361
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:361:
	Glu Gln Gln Pro Glu Thr Lys Pro Gly Lys Gly Thr Leu Gly Lys Gln 1 5 10 15
40	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser
	(362) INFORMATION FOR SEQ ID NO:362
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
. 50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:362:

5	1 5 10 15 15 15 15 15 15 15 15 15 15 15 15 15
	Lys Lys Asn Lys Ser Ser Thr Pro Glu Phe 20 25
10	(363) INFORMATION FOR SEQ ID NO:363
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:363:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
	(364) INFORMATION FOR SEQ ID NO:364
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:364:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
40	Lys Gln Gly Thr Ser Thr Thr Ser Glu Thr
	(365) INFORMATION FOR SEQ ID NO:365
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:365:

5	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
5	
	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
10	(366) INFORMATION FOR SEQ ID NO:366
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:366:
20	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
	Lys Gln Asp Thr Ser Ser Thr Ser Glu Thr
25	(367) INFORMATION FOR SEQ ID NO:367
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:367:
35	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
40	Lys Gln Gly Thr Ser Ser Thr Ser Gly Thr
•	(368) INFORMATION FOR SEQ ID NO:368
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:368:

5	1 5 10 15
	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
10	(369) INFORMATION FOR SEQ ID NO:369
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:369:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr 20 25
	(370) INFORMATION FOR SEQ ID NO:370
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:370:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
40	Lys Gln Ile Thr Ser Ser Thr Ser Glu Thr 20 25
	(371) INFORMATION FOR SEQ ID NO:371
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
-	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:371:

5	1 5 10 15
	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr 20 25
10	(372) INFORMATION FOR SEQ ID NO:372
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:372:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Arg Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys His Val Thr Ser Ser Thr Ser Glu Thr 20 25
	(373) INFORMATION FOR SEQ ID NO:373
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:373:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Thr Glu Gln Pro 1 5 10 15
40	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr
	(374) INFORMATION FOR SEQ ID NO:374
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
E0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:374:

5	Giu Gin Gin Ala Glu Leu Lys Pro Gly Lys Gly Asn Thr Glu Gln Pro 1 5 10 15
	Lys Leu Ile Thr Ser Ser Thr Ser Glu Thr 20 25
10	(375) INFORMATION FOR SEQ ID NO:375
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(X1) SEQUENCE DESCRIPTION: SEQ ID NO:375:
20	Thr Gly Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Glu Gln Gly 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Asp Arg 20 25
	(376) INFORMATION FOR SEQ ID NO:376
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:376:
	Gln Tyr Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Arg Gln Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(377) INFORMATION FOR SEQ ID NO:377
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:377:

- ·	Gln Gln Gln Ala Val Leu Arg His Gly Lys Gly Thr His Gly Gln Glu 1 5 10 15
5	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(378) INFORMATION FOR SEQ ID NO:378
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:378:
20	Gln Gln Gln Thr Lys Leu Gly Pro Gly Arg Gly Thr Pro Gly Gln Gly 1 5 10 15
25	Arg Lys Gly Lys Ser Ser Thr Ser Gly Ser 20 25
	(379) INFORMATION FOR SEQ ID NO:379
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:379:
	Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Val Tyr Phe Ala 20 25
	(380) INFORMATION FOR SEQ ID NO:380
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:380:

Glu Gln Gln Ala Glu Leu Lys Ala Gly Lys Gly Thr Pro Gly Gln Gln 5 Lys Gln Gly Glu Ser Thr Arg Ser Glu Thr 20 10 (381) INFORMATION FOR SEQ ID NO:381 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
(B) TYPE: amino acid
(C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:381: Gln Gln Lys Ala Glu Leu Ala Ala Ser Lys Gly Thr Pro Gly Gln Glu 20 Lys Lys Gly Arg Ser Ser Thr Ser Glu Ser 20 25 (382) INFORMATION FOR SEQ ID NO:382 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids(B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:382: 35 Gln Gln Gln Thr Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu Lys Arg Gly Lys Ser Ser Asn Leu Arg Leu 20 (383) INFORMATION FOR SEQ ID NO:383 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 45 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:383:

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5	1 5 10 15
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 20 25
10	(384) INFORMATION FOR SEQ ID NO:384
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15 ·	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:384:
20	Glu Gln Gln Ala Asp Leu Lys Leu Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Leu Ala Thr Pro Ser Thr Ser Glu Thr 20 25
	(385) INFORMATION FOR SEQ ID NO:385
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:385:
	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Asp Lys Ser 1 5 10 15
40	Asp Val Lys Asp Asn Ala Lys Ser Glu Thr
	(386) INFORMATION FOR SEQ ID NO:386
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:386:

5	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly His Pro 1 5 10 15
,	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(387) INFORMATION FOR SEQ ID NO:387
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:387:
20	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
	(388) INFORMATION FOR SEQ ID NO:388
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:388:
	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr
	(389) INFORMATION FOR SEQ ID NO:389
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:389:

5	1 5 10 15
•	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(390) INFORMATION FOR SEQ ID NO:390
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:390:
20	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Asn Thr Ser Lys Thr Thr Glu Thr 20 25
	(391) INFORMATION FOR SEQ ID NO:391
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:391:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asp Pro 1 5 10 15
40	Ser Lys Thr Thr Ser Lys Thr Thr Glu Thr
	(392) INFORMATION FOR SEQ ID NO:392
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:392:

5	1 5 10 15
	Ser Lys Thr Thr Ser Lys Thr Thr Glu Thr 20 25
10	(393) INFORMATION FOR SEQ ID NO:393
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:393:
20	Asp His Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
. 25	Ser Lys Asn Thr Ser Lys Thr Thr Glu Thr 20 25
	(394) INFORMATION FOR SEQ ID NO:394
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:394:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Arg Ser Thr Ser Lys Thr Thr Glu Thr
	(395) INFORMATION FOR SEQ ID NO:395
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:395:

5	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ala Gly Ser Pro Gly Asn Pro 1 5 10 15
	Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr 20 25
10	(396) INFORMATION FOR SEQ ID NO:396
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:396:
20	Glu Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Ser Glu Thr 20 25
	(397) INFORMATION FOR SEQ ID NO:397
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:397:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Asn Thr Ser Lys Thr Ile Glu Thr
	(398) INFORMATION FOR SEQ ID NO:398
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:398:

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5	Asp Gin Gin Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asp Pro 1 5 10 15
	Ser Lys Asn Thr Ser Lys Thr Pro Glu Thr 20 25
10	(399) INFORMATION FOR SEQ ID NO:399
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:399:
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
	(400) INFORMATION FOR SEQ ID NO:400
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:400:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Asn Thr Ser Glu Thr Thr Glu Thr
	(401) INFORMATION FOR SEQ ID NO:401
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:401:

5	1 5 10 15
	Ser Lys Asn Thr Ser Glu Thr Thr Glx Thr 20 25
10	(402) INFORMATION FOR SEQ ID NO:402
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
٠.	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:402:
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Ser Glu Thr 20 25
	(403) INFORMATION FOR SEQ ID NO:403
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:403:
	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Arg Thr Thr Glu Thr
	(404) INFORMATION FOR SEQ ID NO:404
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:404:

Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 5 Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr 10 (405) INFORMATION FOR SEQ ID NO:405 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:405: 20 Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Phe Pro Gly Asn Pro Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25 (406) INFORMATION FOR SEQ ID NO:406 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids 30 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 406: 35 Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro Ser Lys Ser Thr Ser Lys Thr Asn Glu Thr 40 _ _ 20 (407) INFORMATION FOR SEQ ID NO:407 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:407:

5	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly 1 5 10	Asn Pro 15
	Ser Lys Ser Thr Phe Lys Thr Ser Glu Thr	.*
10	(408) INFORMATION FOR SEQ ID NO:408	
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:408:	
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly 1 5 10	Asn Pro 15
25	Ser Lys Ser Thr Ser Thr Thr Ser Glu Thr 20 25	·
	(409) INFORMATION FOR SEQ ID NO:409	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:409:	
	Glu Gln Gln Leu Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly 1 5 10	Asn Pro
4 0	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr	
	(410) INFORMATION FOR SEQ ID NO:410	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
50	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:410:	

5	Gln Gln Gln Pro Gly Leu Lys Pro Ser Phe Gly Pro Pro Gly Lys Pro 1 15
	Ser Gln Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(411) INFORMATION FOR SEQ ID NO:411
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:411:
20	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
25	Thr Lys Ser Asn Ser Lys Gln Thr Asp Thr 20 25
20	(412) INFORMATION FOR SEQ ID NO:412
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:412:
•	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
40	Ala Lys Ser Asn Ser Lys Gln Thr Asp Thr 20 25
	(413) INFORMATION FOR SEQ ID NO:413
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:413:
	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 10 15

5	20 25
	(414) INFORMATION FOR SEQ ID NO:414
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:414:
	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
20	Ala Ile Ser Asn Ser Lys Gln Thr Asp Thr 20 25
	(415) INFORMATION FOR SEQ ID NO:415
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:415:
35	Gln Gln Lys Pro Gly Leu Gln Pro Ser Ser Gly Ser Pro Gly Lys Ala 1 5 10 15
35	Ala Ile Ser Asn Ser Lys Gln Ser Asn Thr
	20 25
40	(416) INFORMATION FOR SEQ ID NO:416
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:416:
50	Gln Gln Lys Pro Gly Leu Gln Pro Ser Ser Gly Ser Pro Gly Lys Ala 1 5 10 15
	Ala Ile Ser Asn Ser Lys Gln Ala Asn Thr 20 25
55	

	(417) INFORMATION FOR SEQ ID NO:417
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:417:
15	Gln Gln Lys Pro Val Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 . 15
	Ala Met Ser Asn Ser Lys Gln Ile Asp Thr 20 25
20	(418) INFORMATION FOR SEQ ID NO:418
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:418:
30	Gln Gln Lys Pro Ser Leu Gln Pro Ser Ser Asp Ser Pro Gly Lys Ala 1 5 10 15
35	Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25
	(419) INFORMATION FOR SEQ ID NO:419
4 0	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:419:
	Glu Arg Val Gly Asp Leu Glu Pro Gly Arg Gly Ile Pro Gly Lys Ala 1 5 10 15
50	Pro Lys Gly Asp Ser Lys Lys Ile Glu Thr 20 25
	(420) INFORMATION FOR SEQ ID NO:420

5	(1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:420:
	Glu Arg Val Gly Asp Leu Glu Pro Glu Arg Gly Ile Pro Gly Lys Ala 1 5 10 15
15	Pro Lys Gly Asp Ser Lys Lys Ile Glu Thr 20 25
	(421) INFORMATION FOR SEQ ID NO:421
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:421:
30	Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
30	Pro Lys Gly Asp Ser Lys Lys Thr Glu Thr 20 25
35	(422) INFORMATION FOR SEQ ID NO:422
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(<u>i</u> i) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:422:
45	Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Gly Lys Ala 1 5 10 15
50	Ser Lys Gly Asp Ser Lys Lys Thr Glu Thr 20 25
	(423) INFORMATION FOR SEQ ID NO:423
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
55	(,

	(C) TOPOLOGY: linear
5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:423:
10	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asx 1 5 10 15
15	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 25
	(424) INFORMATION FOR SEQ ID NO:424
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:424:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
30	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(425) INFORMATION FOR SEQ ID NO:425
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 425:
45	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Arg Gly Asn Ser Lys Arg Ala Glu Thr 20 25
50	(426) INFORMATION FOR SEQ ID NO:426
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid
55	(C) TOPOLOGY: linear

	(ii) MOLECULE TYPE: peptide
5	(X1) SEQUENCE DESCRIPTION: SEQ ID NO:426:
. 10	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
45	(427) INFORMATION FOR SEQ ID NO:427
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:427:
25	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Lys Arg Ala Glu Thr 20 25
30	(428) INFORMATION FOR SEQ ID NO:428
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:428:
40	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 10 15
45	Ser Lys Gly Asp Ser Arg Arg Ala Glu Thr 20 25
	(429) INFORMATION FOR SEQ ID NO:429
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 429:
5	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 10 15
10	Ser Lys Gly Asn Ser Arg Arg Ala Glu Thr 20 25
	(430) INFORMATION FOR SEQ ID NO:430
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:430:
	Gln Gln Val Gly Gly Leu Glu Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
25	Ser Lys Gly Asx Ser Lys Arg Ala Glu Thr 20 25
	(431) INFORMATION FOR SEQ ID NO:431
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:431:
40	Glu Gln Leu Gly Asp Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
45	(432) INFORMATION FOR SEQ ID NO:432
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:432:

5	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Thr Gly Lys Asp 1 10 15
	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 25
10	(433) INFORMATION FOR SEQ ID NO:433
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:433:
	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(434) INFORMATION FOR SEQ ID NO:434
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:434:
	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Ile Pro Gly Lys Asp 1 5 10 15
4 0	Ser Lys Gly Asn Ser Lys Arg Pro Glu Thr
	(435) INFORMATION FOR SEQ ID NO:435
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:435:

5	Glu Gln Val Gly Gly Val Gln Pro Gly Arg Gly Ile Pro Gly Lys Asp 1 10 15
	Ser Lys Gly Asp Ser Lys Arg Pro Glu Thr 20 25
	(436) INFORMATION FOR SEQ ID NO:436
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
15	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:436:
20	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Asn Gly Asp Ser Lys Arg Pro Glu Thr 20 25
25	(437) INFORMATION FOR SEQ ID NO:437
	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 26 amino acids (B) TYPE: amino acid
30	(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:437:
33	Gln Lys Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 10 15
40	Ser Lys Gly Asn Ser Lys Arg Thr Glu Thr 20 25
	(438) INFORMATION FOR SEQ ID NO:438
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
40	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:438:
	Gln Glu Val Gly Gly Val Glx Pro Gly Arg Gly Thr Pro Gly Lys Asx 1 5 10 15
55	

 5.	Ser Lys Gly Asx Ser Lys Arg Ala Glu Thr 20 25
	(439) INFORMATION FOR SEQ ID NO:439
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:439:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
20	Ser Asn Gly Asp Ser Lys Gln Ala Glx Thr 20 25
•	(440) INFORMATION FOR SEQ ID NO:440
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:440:
35	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Ser Pro Gly Lys Asp 1 5 10 15
	Thr Asn Gly Asp Ser Lys Glu Ala Glx Thr 20 25
40	(441) INFORMATION FOR SEQ ID NO:441
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
4 5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:441:
50	Ala Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Asn Gly Asp Ser Lys Gln Ala Glx Ser 20 25
55	

	(442) INFORMATION FOR SEQ ID NO:442
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(X1) SEQUENCE DESCRIPTION: SEQ ID NO:442:
15	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val 1 5 10 15
	Ser Gln Gly Asp Ser Lys Gln Ala Glx Thr 20 25
20	(443) INFORMATION FOR SEQ ID NO:443
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:443:
30	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val 1 5 10 15
35	Ser Gln Gly Asp Ser Lys Glu Pro Glx Thr 20 25
	(444) INFORMATION FOR SEQ ID NO:444
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
-	(ii) MOLECULE TYPE: peptide
4 5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:444:
	Glu Gln Leu Gly Gly Leu Gln Pro Glu Arg Gly Thr Pro Gly Lys Glu 1 5 10 15
50	Ser Lys Gly Asn Ser Met Arg Ala Glu Thr 20 25
	(445) INFORMATION FOR SEQ ID NO:445

5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:445:
	Glu Gln Val Gly Asp Leu Gln Pro Gly Arg Gly Asx Pro Gly Lys Asp 1 10 15
15	Ser Lys Gly Asn Ala Lys Arg Val Glu Thr 20 25
	(446) INFORMATION FOR SEQ ID NO:446
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:446:
30	Glu Gln Val Gly Asp Leu Gln Pro Gly Arg Gly Asn Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Gln Arg Pro Glu Thr 20 25
35	(447) INFORMATION FOR SEQ ID NO:447
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) _MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:447:
45	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Leu Gly Lys Asp 1 5 10 15
50	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(448) INFORMATION FOR SEQ ID NO:448
55	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids

(B) TYPE: amino acid

_	(C) TOPOLOGY: linear
5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:448:
10	Gln Glx Val Gly Gly Ala Glx Pro Gly Arg Gly Ser Pro Gly Lys Ala 1 5 10 15
15	Ser Lys Gly Asx Ser Lys Arg Ala Glu Thr 20 25
	(449) INFORMATION FOR SEQ ID NO:449
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:449:
	Gln Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Ser Pro Gly Lys Asp 1 5 10 15
30	Ser Lys Gly Asn Ala Gln Arg Thr Glx Thr 20 25
	(450) INFORMATION FOR SEQ ID NO:450
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:450:
45	Asp Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Gly Lys Asn 1 5 10 15
	Ser Asn Gly Asp Ser Lys Thr Pro Glx Thr 20 25
50	(451) INFORMATION FOR SEQ ID NO:451
55	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
,,,	(-/

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:451:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Ser Arg Glu Asp 1 5 10 15
10	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(452) INFORMATION FOR SEQ ID NO:452
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:452:
25	Glu Gln Val Gly Ala Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 10 15
	Ser Gln Ala Asp Ser Lys Glu Ala Glx Thr 20 25
30	(453) INFORMATION FOR SEQ ID NO:453
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 22 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:453:
4 0 -	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val
45	Glu Gly Ser Val Glu Thr 20
	(454) INFORMATION FOR SEQ ID NO:454
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide

	(X1) SEQUENCE DESCRIPTION: SEQ ID NO:454:
5	Glu Gln Val Gly Ala Phe Gln Pro Gly Arg Gly Asn Ser Gly Lys Ala 1 5 10 15
10	Ser Lys Gly Asp Ser Lys Arg Pro Asp Thr 20 25
	(455) INFORMATION FOR SEQ ID NO:455
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:455:
	Glu Gln Val Gly Ala Phe Gln Pro Gly Lys Gly Asn Ser Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Asp Ser Lys Arg Pro Asp Thr 20 25
	(456) INFORMATION FOR SEQ ID NO:456
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:456:
40	Glu Gln Val Gly Ala Phe Gln Pro Gly Lys Gly Asn Ser Gly Lys Ala 1 5 10 15
	Ser Lys Gly Asp Ser Asn Arg Pro Asp Thr 20 25
45	(457) INFORMATION FOR SEQ ID NO:457
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:457:

5	Gln Gln Val Gly Gly Val Gln Ala Gly Arg Ala Asn Pro Gly Lys Asp 1 5 10 15
	Ser Arg Gly Ile Ser Lys Arg Thr Glu Thr 20 25
10	(458) INFORMATION FOR SEQ ID NO:458
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:458:
20	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Gln Gly Glu Ser Thr Arg Ser Glu Thr 20 25
	(459) INFORMATION FOR SEQ ID NO:459
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:459:
	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Gln Gly Thr Ser Thr Arg Ser Glu Thr 20 25
	(460) INFORMATION FOR SEQ ID NO:460
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:460:

5	1 5 10 Lys Pro Gly Lys Gly Thr Pro Gly Gln Gl	n
	Lys Gln Gly Thr Ser Ala Arg Ser Glu Thr 20 25	
	(461) INFORMATION FOR SEQ ID NO:461	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
15	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:461:	
20	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gl 1 5 10 15	n
	Lys Gln Gly Thr Ser Ile Arg Ser Asp Thr 20 25	
25	(462) INFORMATION FOR SEQ ID NO:462	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:462:	
35	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gl 1 5 10 15	u
40	Lys Gln Gly Thr Ser Ile Arg Ser Asp Thr 20 25	
	(463) INFORMATION FOR SEQ ID NO:463	
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:463:	
	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gl 1 5 10 15	n

	Asn Gln Gly Thr Ser Thr Arg Ser Asp Thr 20 25
5	(464) INFORMATION FOR SEQ ID NO:464
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 464:
15	Gln Gln Val Gly Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Gln Gln 1 5 10 15
20	Lys Gln Asp Thr Ser Thr Arg Ser Asp Thr 20 25
	(465) INFORMATION FOR SEQ ID NO:465
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:465:
	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly His Pro 1 5 10 15
35	Arg Gln Gly Ala Ser Phe Arg Ser Asp Ser 20 25
	(466) INFORMATION FOR SEQ ID NO:466
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
4 5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:466:
50	Gln Gln Val Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glr 1 5 10 15
	Gly Thr Gly Thr Ser Val Lys Ala Glu Thr 20 25
55	

	(467) INFORMATION FOR SEQ ID NO:467
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:467:
15	Glu Gln Val Ala Glu Val Lys Pro Gly Lys Gly Ser Pro Gly Lys Pro 1 5 10 15
•	Ser Gln Gly Lys Ser Ile Lys Ala Ser Thr 20 25
20	(468) INFORMATION FOR SEQ ID NO:468
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:468:
30	Glu Gln Val Ala Glu Val Lys Pro Gly Arg Gly Ser Pro Gly Lys Pro 1 5 10 15
35	Ser Gln Gly Lys Ser Ile Lys Ala Ser Thr 20 25
	(469) INFORMATION FOR SEQ ID NO:469
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
4 5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:469:
	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Asp Pro Gly Arg Pro 1 5 10 15
50	Arg Gln Ala Ser Ser Thr Ile Ser Ala Thr 20 25
	(470) INFORMATION FOR SEQ ID NO:470

5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:470:
	Glu Gln Val Ala Glu Val Pro Gln Gly Lys Gly Arg Pro Gly Lys Ser 1 5 10 15
15	Leu Gln Gly Lys Ser Leu Lys Ala Ser Thr 20 25
	(471) INFORMATION FOR SEQ ID NO:471
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:471:
30	Gln Gln Met Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Lys Pro 1 5 10 15
	Gly Val Val Pro Ser Phe Phe Ser Glu Thr 20 25
35	(472) INFORMATION FOR SEQ ID NO:472
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	(<u>ii</u>) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:472:
45	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Arg Tyr 1 5 10 15
50	Ile Trp Glu Pro Ser Phe Phe Asn Glu Gly 20 25
	(473) INFORMATION FOR SEQ ID NO:473
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids
55	

5	(B) TYPE: amino acid(C) TOPOLOGY: linear
`	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:473:
10	Gln Gln Gln Ala Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
15	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(474) INFORMATION FOR SEQ ID NO:474
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:474:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
30	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(475) INFORMATION FOR SEQ ID NO:475
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:475:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
45	Ser Lys Ser Thr Ser Asn Thr Ala Ala Thr 20 25
	(476) INFORMATION FOR SEQ ID NO:476
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
55	

	(11) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:476:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Ala Gly Lys Pro 1 5 10 15
10	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(477) INFORMATION FOR SEQ ID NO:477
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:477:
25	Arg Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Pro Pro Gly Lys Pro 1 10 15
	Ser Arg Gly Thr Ser Arg Ser Ala Ala Thr 20 25
30	(478) INFORMATION FOR SEQ ID NO:478
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:478:
40	Gln Gln Gln Ala Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Arg Thr 1 _ 5 10 15
4 5	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(479) INFORMATION FOR SEQ ID NO:479
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:479:
5	Gln Gln Glu Pro Gly Leu Arg Pro Ser Ser Gly Thr Pro Gly Arg Thr 1 5 10 15
10	Pro Arg Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(480) INFORMATION FOR SEQ ID NO:480
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:480:
	Xaa Gln Glu Pro Gly Leu Arg Pro Ser Ser Gly Ser Pro Gly Arg Thr 1 5 10 15
25	Pro Arg Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(481) INFORMATION FOR SEQ ID NO:481
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:481:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Ser Arg Val
40	1 5 10 15
	Ser Lys Ser Thr Ser Lys Thr Pro Glu Thr 20 25
45	(482) INFORMATION FOR SEQ ID NO:482
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:482:
55	

5	Gln His Gln Ala Gly Leu Lys Arg Ser Ser Gly Pro Pro Gly Lys Pro 1 5 10 15
	Ser Thr Ser Thr Ser Lys Thr Ala Ala Thr 20 25
10	(483) INFORMATION FOR SEQ ID NO:483
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:483:
20	Glx Gln Glu Ser Gly Leu Lys Pro Thr Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
25	Ser Lys Ser Arg Ser Lys Ala Ala Asp Ala 20 25
	(484) INFORMATION FOR SEQ ID NO:484
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
•	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:484:
	Gln Thr Lys Pro Thr Leu Lys Pro Thr Thr Gly Ser Pro Gly Arg Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Lys Asp Pro Val Thr
	(485) INFORMATION FOR SEQ ID NO:485
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide(xi) SEQUENCE DESCRIPTION: SEQ ID NO:485:

5	1 5 10 15
	Ser Arg Ser Thr Ser Arg Asp Pro Val Ser 20 25
10	(486) INFORMATION FOR SEQ ID NO:486
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:486:
20	Glu Thr Arg Pro Ala Leu Lys Pro Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
	Ser Lys Thr Thr Ser Lys Asp Pro Val Thr 20 25
25	(487) INFORMATION FOR SEQ ID NO:487
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:487:
35	Gln Asn Arg Pro Ala Leu Lys Ala Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
40	Ser Glu Thr Thr Ser Lys Asp Pro Ala Thr 20 25
-	(488) INFORMATION FOR SEQ ID NO:488
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:488:
	Gln Thr Thr Pro Ala Leu Lys Pro Lys Thr Gly Ser Pro Gly Lys Thr 1 5 10 15

5	20 25
-	(489) INFORMATION FOR SEQ ID NO:489
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:489:
.5	Gln Thr Arg Pro Ala Leu Arg Pro Thr Thr Gly Ser Pro Gly Glu Ala 1 5 10 15
20	Ser Glu Thr Thr Ser Lys Gly Pro Gly Thr 20 25
	(490) INFORMATION FOR SEQ ID NO:490
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 26 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:490:
	Gln Thr Arg Pro Ala Leu Lys Pro Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
35	Ser Glu Thr Thr Ser Arg Asp Thr Ala Tyr 20 25
	(491) INFORMATION FOR SEQ ID NO:491
40	(i)_SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:491:
50	Leu Glu Gly Val Gln Leu Trp Gly Gly Arg Gly Ile Ser Arg Lys Tyr 1 5 10 15
	Ala Lys Gly Asn Gly Lys Arg Glu Asp Ser 20 25

	(492) INFORMATION FOR SEQ ID NO:492
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 10 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:492:
	Tyr Asn Asn Pro Gly Asn Gly Tyr Ile Ala 1 5 10
. 15	(493) INFORMATION FOR SEQ ID NO:493
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 10 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:493:
25	Tyr Ile Asn Pro Gly Lys Gly Tyr Leu Ser
20	(494) INFORMATION FOR SEQ ID NO:494
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 11 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:494:
40	Arg Ala Ser Gln Glu Ile Ser Gly Tyr Leu Ser 1 5 10
	(495) INFORMATION FOR SEQ ID NO:495
4 5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 11 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:495:
55	Arg Ala Ser Gly Asn Ile His Asn Tyr Leu Ala 1 5 10

	(496) INFORMATION FOR SEQ ID NO:496
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 11 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:496:
	Arg Ala Ser Gln Asp Ile Asn Asn Phe Leu Asn 1 5 10
, 15	(497) INFORMATION FOR SEQ ID NO:497
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 11 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
•	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:497:
20	Arg Ala Ser Gln Ser Ile Gly Asn Asn Leu His 1 5 10
30	(498) INFORMATION FOR SEQ ID NO:498
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 7 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
33	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:498:
40	Ala Ala Ser Thr Leu Asp Ser 1 5
	(499) INFORMATION FOR SEQ ID NO:499
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 7 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:499:
55	Tyr Thr Thr Leu Ala Asp 1 5

	(500) INFORMATION FOR SEQ ID NO:500
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 7 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:500:
	Phe Thr Ser Arg Ser Gln Ser 1 5
15	(501) INFORMATION FOR SEQ ID NO:501
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 7 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:501:
	Lys Ala Ser Ser Leu Glu Ser 1 5
30	(502) INFORMATION FOR SEQ ID NO:502
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 9 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:502:
40	Leu Gln Tyr Leu Ser Tyr Pro Leu Thr 1 5
	(503) INFORMATION FOR SEQ ID NO:503
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 9 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:503:
	Gln His Phe Trp Ser Thr Pro Arg Thr

	(504) INFORMATION FOR SEQ ID NO:504
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 9 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:504:
	Gln Gln Gly Asn Ala Leu Pro Arg Thr
15	(505) INFORMATION FOR SEQ ID NO:505
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 7 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:505:
	Gln Gln Tyr Asn Ser Tyr Ser 1 5
30	(506) INFORMATION FOR SEQ ID NO:506
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 5 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:506:
40	Thr Phe Gly Ile Thr 1 5
	(507) INFORMATION FOR SEQ ID NO:507
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:507:
	Gly Tyr Gly Val Asn
55	1 5

	(508) INFORMATION FOR SEQ ID NO:508
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 5 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:508:
	Ser Asn Gly Ile Asn 1 5
15	(509) INFORMATION FOR SEQ ID NO:509
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid
٠.	(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:509:
	Asp Tyr Ala Met His 1 5
30	(510) INFORMATION FOR SEQ ID NO:510
35	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 10 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:510:
40	·
•	Glu Ile Phe Pro Gly Asn Ser Lys Thr Tyr 1 5 10
45	1 5 10 (511) INFORMATION FOR SEQ ID NO:511 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid
45	1 5 10 (511) INFORMATION FOR SEQ ID NO:511 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	1 5 10 (511) INFORMATION FOR SEQ ID NO:511 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide
	1 5 10 (511) INFORMATION FOR SEQ ID NO:511 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

	(512) INFORMATION FOR SEQ ID NO:512
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 10 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:512:
	Tyr Asn Asn Pro Gly Asn Gly Tyr Ile Ala 1 5 10
15	(513) INFORMATION FOR SEQ ID NO:513
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:513:
	Ile Ser Trp Asp Ser Ser Ser Ile Gly 1 5
30	(514) INFORMATION FOR SEQ ID NO:514
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 5 amino acids(B) TYPE: amino acid
35	(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:514:
40	Arg Glu Ile Arg Tyr 1 5
	(515) INFORMATION FOR SEQ ID NO:515
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 8 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:515:
55	Glu Arg Asp Tyr Arg Leu Asp Tyr 1 5

	(516) INFORMATION FOR SEQ ID NO:516
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 12 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:516:
15	Ser Glu Tyr Tyr Gly Gly Ser Tyr Lys Phe Asp Tyr 1 5 10
	(517) INFORMATION FOR SEQ ID NO:517
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 17 amino acids(B) TYPE: amino acid(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:517:
	Gly Arg Asp Tyr Tyr Asp Ser Gly Gly Tyr Phe Thr Val Ala Phe Asp 1 5 10 15
30	Ile
	(518) INFORMATION FOR SEQ ID NO:518
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 11 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:518:
45	Arg Ala Ser Gln Ser Ile Ser Arg Trp Leu Ala 1 5 10
4 5	(519) INFORMATION FOR SEQ ID NO:519
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 7 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
EE	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:519:

Glu Ala Ser Asn Asp Leu Ala (520) INFORMATION FOR SEQ ID NO:520 5 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 10 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:520: 15 Asp Phe Tyr Met Glu (521) INFORMATION FOR SEQ ID NO:521 20 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 9 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 25 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:521: Ile Ile Trp Asp Asp Gly Ser Asp Gln 30 1 (522) INFORMATION FOR SEQ ID NO:522 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 11 amino acids 35 (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:522: Gln Ala Ser Gln Ser Ile Ile Lys Tyr Leu Asn 45

Claims

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 A method for determining how to humanize a rodent antibody or fragment thereof by resurfacing, said method comprising:

(a) determining the conformational structure of the variable region of said rodent antibody or fragment thereof by constructing a three-dimensional model of said rodent antibody variable region;

(b) generating sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of heavy and light chain framework positions wherein said set is identical in 98% of said sufficient number of rodent antibody heavy and light chains;

- (c) defining for said rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using said set of framework positions generated in said step (b); (d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c), wherein said heavy and light chain from said human antibody are or are not naturally paired;
- (e) substituting, in the amino acid sequence of said rodent antibody or fragment thereof to be humanized said set of heavy and light chain surface exposed amino acid residues defined in said step (c) with said set of heavy and light chain surface exposed amino acid residues identified in said step (d);
- (f) constructing a three-dimensional model of said variable region of said rodent antibody or fragment thereof resulting from the substituting specified in said step (e);
- (g) identifying, by comparing said three-dimensional models constructed in said steps (a) and (f), any amino acid residues from said set identified in said step (d), that are within 5 Angstroms of any atom of any residue of the complementarity determining regions of said rodent antibody or fragment thereof to be humanized; and
- (h) changing any residues identified in said step (g) from the human to the original rodent amino acid residue to thereby define a rodent antibody humanizing set of surface exposed amino acid residues; with the proviso that said step (a) need not be conducted first, but must be conducted prior to said step (g).
- 2. The method of claim 1, wherein said rodent antibody is an antibody fragment.

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- 3. The method of claim 2, wherein said rodent antibody fragment is a single chain antibody, a F_V fragment, a Fab fragment, a Fab₂ fragment or a Fab' fragment.
- 4. The method of claim 1 or 2, wherein said step (d) identifies a set of naturally paired heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c).
- 5. The method of claim 1 or 2, wherein said surface exposed amino acid residues are those residues whose solvent accessibility is above 30%.
 - The method of claim 1 or 2, wherein the rodent antibody or fragment thereof to be humanized is a murine antibody.
- 7. The method of claim 6, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

	Light Chain		
40	Position	Human	Mouse
	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
45	5	T 61 L 37	Т 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
	15	P 62 V, 25 L 12	L 47 P 30 V 8 A 7
50	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
	46	P 94	P 82 S 9
	47	G 89	G 71 D 18
55	51	K 43 R 31	K 70 Q 13 R 8 T 5
	63	G 91	G 98

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	66	D 43 S 25 A 9	D 38 A 26 S 26
	73	S 96	S 90 I 5
5	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
•	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
,	87	E 71 D 11 G 7	E 91 D 6
10	111	K 74 R 12 N 6	K 93
	115	K 54 L 40	K 87 L 5
	116	R 60 G 33 S 5	R 89 G 9
15	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
		Heavy Chain	
	Position	Human	Mouse
20	118	E 47 Q 46	E 59 Q 29 D 10
	120	Q 83 T 7	Q 68 K 26
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5
25	126	G 54 A 23 P 18	G 36 P 30 A 29
	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
	128	L 61 V 31 F 7	L 96
30	130	K 46 Q 41 E 5	K 52 Q 27 R 17
	131	P 95	P 91 A 5
	132	G 74 S 16 T 7	G 82 S 17
35	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
	143	G 96	G 98
40	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
40	160	P 84 S 10	P 89 H 7
	161	G 93	G 71 E 24
45	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
	184	S 70 K 9 P 8	K 42 S 37 T 6
50	186	K 53 Q 22 R 7 N 5	K 83 Q 7
•	187	G 66 S 21 T 5	G 62 S 18 D 10
	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
55	196	S 91	S 76 A 16
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11
	1	1	ı

· Table 1				
212	T 91			
210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43		
209	A 50 P 21 S 13 T 8	E 88 D 7		
208	R 46 T 18 K 17 D 6	S 67 A 14 T 11		

10

- The method of claim 1 or 2, wherein the rodent antibody or fragment thereof to be humanized is murine antibody anti-N901.
- 9. The method of claim 8, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

	Light Chain		
	Position	Human	Mouse
20 .	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
• • •	5	T 61 L 37	Т 87
25	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
•	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
30	46	P 94	P 82 S 9
	47	G 89	G 71 D 18
0.5	51	K 43 R 31	K 70 Q 13 R 8 T 5
35	63	G 91	G 98
•	66	D 43 S 25 A 9	D 38 A 26 S 26
40	73	S 96	S 90 I 5
40	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
•	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
45	87	E 71 D 11 G 7	E 91 D 6
	111	K 74 R 12 N 6	K 93
	115	K 54 L 40	K 87 L 5
50	116	R 60 G 33 S 5	R 89 G 9
50	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
		Heavy Chain	
55	Position	Human	Mouse
55	118	E 47 Q 46	E 59 Q 29 D 10

			1
	120	Q 83 T 7	Q 68 K 26
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5
5	126	G 54 A 23 P 18	G 36 P 30 A 29
	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
	128	L 61 V 31 F 7	L 96
10	130	K 46 Q 41 E 5	K 52 Q 27 R 17
	131	P 95	P 91 A 5
	132	G 74 S 16 T 7	G 82 S 17
15	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
	143	G 96	G 98
	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
20	160	P 84 S 10	P 89 H 7 .
	161	G 93	G 71 E 24
25	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
	184	S 70 K 9 P 8	K 42 S 37 T 6
30	186	K 53 Q 22 R 7 N 5	K 83 Q 7
	187	G 66 S 21 T 5	G 62 S 18 D 10
	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
35	196	S 91	S 76 A 16
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11,
	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11
40	209	A 50 P 21 S 13 T 8	E 88 D 7
	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43
	212	T 91	
45		Table 1	

- 10. A method for producing a humanized rodent antibody or fragment thereof from a rodent antibody or fragment thereof by resurfacing, said method comprising.
 - (I) carrying out the method of claim 1; and
 - (II) modifying the rodent antibody or fragment thereof by replacing the set of rodent antibody surface exposed amino acid residues with the rodent antibody humanizing set of surface exposed amino acid residues defined in said step (h).
 - 11. The method of claim 10, wherein said rodent antibody is an antibody fragment.
 - 12. The method of claim 11, wherein said rodent antibody fragment is a single chain antibody, a F_V fragment,

a Fab fragment, a Fab₂ fragment or a Fab' fragment.

- 13. The method of claim 10 or 11, wherein said step (d) identifies a set of naturally paired heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c).
- 14. The method of claim 10 or 11, wherein said surface exposed amino acid residues are those residues whose solvent accessibility is above 30%.
- 15. The method of claim 10 or 11, wherein the rodent antibody or fragment thereof to be humanized is a murine antibody.
 - **16.** The method of claim 15, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

15	Light Chain		
	Position	Human	Mouse
	1 ·	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
20	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
	5	T 61 L 37	Т 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
25	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
•	46	P 94	P 82 S 9
30	47	G 89	G 71 D 18
	51	K 43 R 31	K 70 Q 13 R 8 T 5
	63	G 91	G 98
35	66	D 43 S 25 A 9	D 38 A 26 S 26
	73	S 96	S 90 I 5
•	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
40	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
	87	E 71 D 11 G 7	E 91 D 6
	111	K 74 R 12 N 6	K 93
45	115	K 54 L 40	K 87 L 5
	116	R 60 G 33 S 5	R 89 G 9
	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
50	Heavy Chain		
·	Position	Human	Mouse
	118	E 47 Q 46	E 59 Q 29 D 10
55 ·	120	Q 83 T 7	Q 68 K 26
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5

•			
	126	G 54 A 23 P 18	G 36 P 30 A 29
	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
5	128	L 61 V 31 F 7	L 96
	130	K 46 Q 41 E 5	K 52 Q 27 R 17
	131	P 95	P 91 A 5
10	132	G 74 S 16 T 7	G 82 S 17
	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
	143	G 96	G 98
15	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
	160	P 84 S 10	P 89 H 7
	161	G 93	G 71 E 24
20	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
25	184	S 70 K 9 P 8	K 42 S 37 T 6
	186	K 53 Q 22 R 7 N 5	K 83 Q 7
	187	G 66 S 21 T 5	G 62 S 18 D 10
30	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
	196	S 91	S 76 A 16
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11
35	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11
	209	A 50 P 21 S 13 T 8	E 88 D 7
	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43
	212	Т 91	-
	Table 1		

17. The method of claim 10 or 11, wherein the rodent antibody or fragment thereof to be humanized is murine antibody anti-N901.

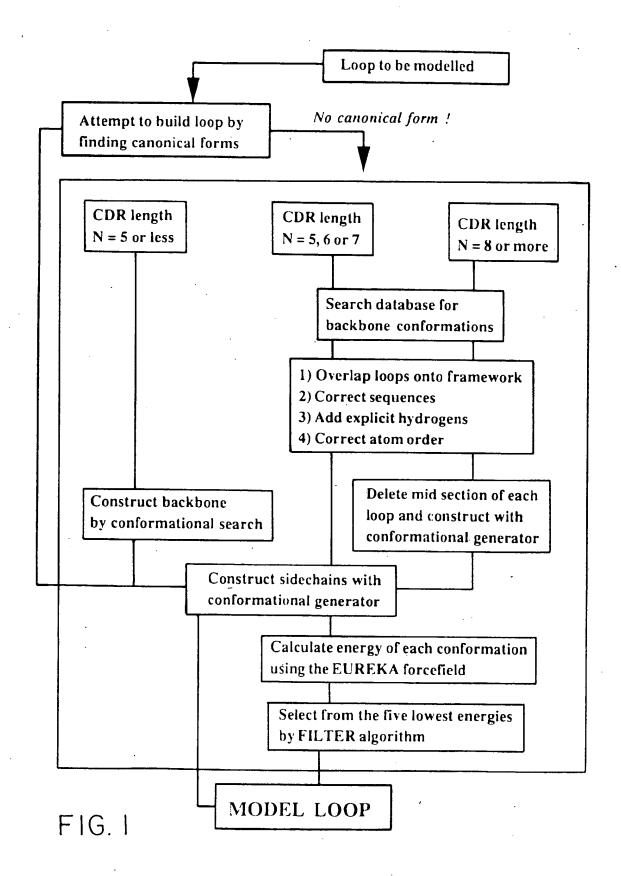
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18. The method of claim 17, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

Light Chain			
Position	Human	Mouse	
1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6	
3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5	

	1		
	5	T 61 L 37	Т 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
5	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
	46	P 94	P 82 S 9
10	47	G 89	G 71 D 18
	51	K 43 R 31	K 70 Q 13 R 8 T 5
•	63	G 91	G 98
5	66	D 43 S 25 A 9	D 38 A 26 S 26
	73	S 96	S 90 I 5
	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
o	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
	87	E 71 D 11 G 7	E 91 D 6
•	111	K 74 R 12 N 6	K 93
5	115	K 54 L 40	K 87 L 5
	116	R 60 G 33 S 5	R 89 G 9
•	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
30 ·		Heavy Chain	-
	Position	Human	
	Position	Trainair	Mouse
	118	E 47 Q 46	E 59 Q 29 D 10
5			
5	118	E 47 Q 46	E 59 Q 29 D 10
5	118	E 47 Q 46 Q 83 T 7	E 59 Q 29 D 10 Q 68 K 26
	118 120 122	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5
	118 120 122 126	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29
	118 120 122 126 127	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6
0	118 120 122 126 127 128	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96
O	118 120 122 126 127 128 130	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17
5	118 120 122 126 127 128 130	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5 P 95	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17 P 91 A 5
5 0	118 120 122 126 127 128 130 131	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5 P 95 G 74 S 16 T 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17 P 91 A 5 G 82 S 17
o 5	118 120 122 126 127 128 130 131 132	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5 P 95 G 74 S 16 T 7 R 53 K 23 S 17 T 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17 P 91 A 5 G 82 S 17 K 66 S 17 R 13
o 5	118 120 122 126 127 128 130 131 132 136 143	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5 P 95 G 74 S 16 T 7 R 53 K 23 S 17 T 7 G 96	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17 P 91 A 5 G 82 S 17 K 66 S 17 R 13 G 98 T 63 S 19 N 7 A 5 D 5
5	118 120 122 126 127 128 130 131 132 136	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18 G 53 E 22 A 14 D 7 L 61 V 31 F 7 K 46 Q 41 E 5 P 95 G 74 S 16 T 7 R 53 K 23 S 17 T 7 G 96 T 46 S 32 N 9 I 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6 L 96 K 52 Q 27 R 17 P 91 A 5 G 82 S 17 K 66 S 17 R 13 G 98

		Table 1		
20	212	T 91		
,	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43	
	209	A 50 P 21 S 13 T 8	E 88 D 7	
15	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11	
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11	
	196	S 91	S 76 A 16	
10	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6	
	187	G 66 S 21 T 5	G 62 S 18 D 10	
	186	K 53 Q 22 R 7 N 5	K 83 Q 7	
5	184	S 70 K 9 P 8	K 42 S 37 T 6	
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11	
	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5	
*	1		1	



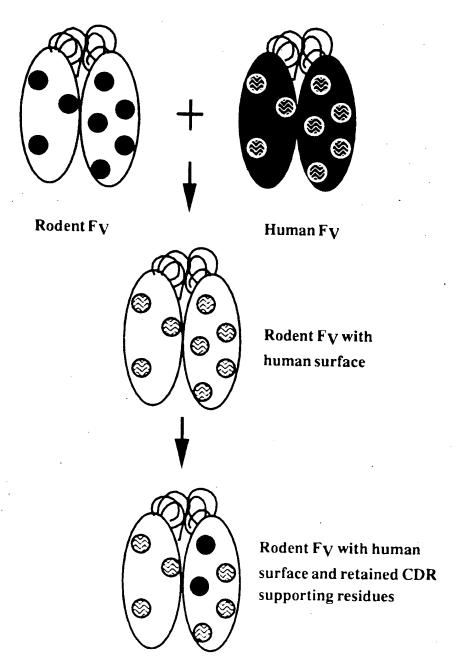
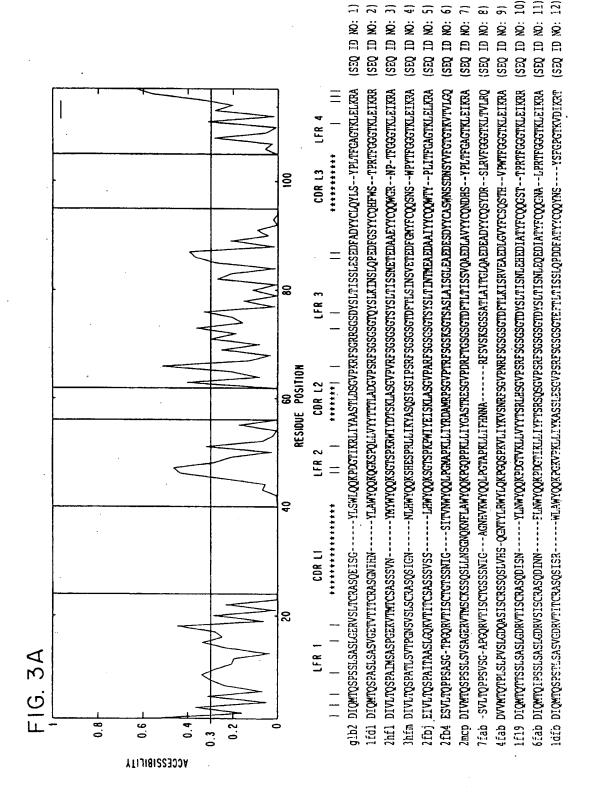


FIG. 2



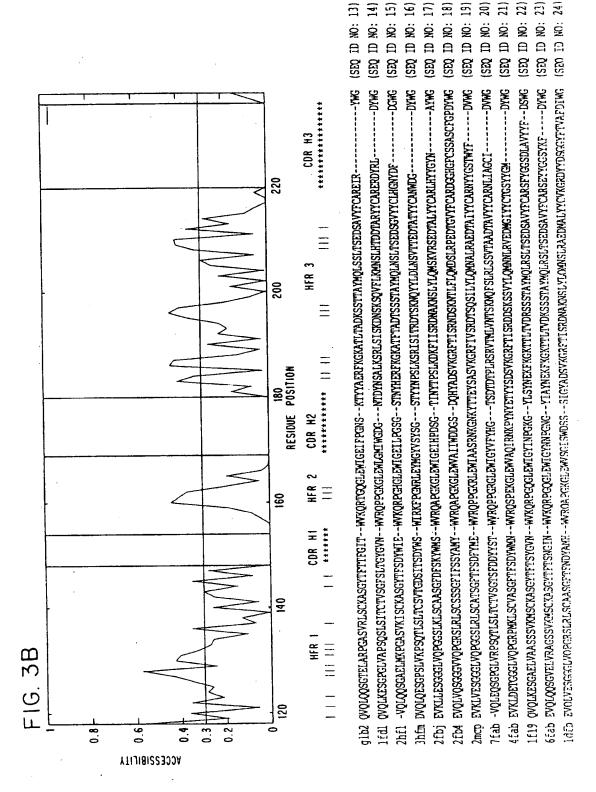


FIG. 4A Light Chain Sequences

·			30 ,	40	50	09	70
N901L		:DVLMTQTPLSLPVSLGDQASISC R	RSSQIIIHSDGNTY-LE		WFLQKPGQSPKLLIY	KVSNRFS	GVPDRFSG
		:QSVLTQPPSASG-TPGQRVTISC S	SGTSSNIGSSTVN WYQQLPGMAPKLLIY	STVN WYQ	QLPGMAPKLLIY	RDAMRPS	GVPDRFSG
3 N901L/KÖL		QVLMTQTPSSLPVTLGQQASISC R	RSSQIIIHSDGNTY-LE	Y-LE WFL	WFLQKPGQSPKLLIY	KVSNRFS	GVPDRFSG
KV2F\$HUMAN	50	: DVVMTQSPLSLPVTLGQPASISC R	RSSQSLVYSDGNTY-LN WFQQRPGQSPRRLIY	Y-LN WFQ	2RPGQSPRRLIY	KVSNRDS	GVPDRFSG
֓֡֓֞֜֜֞֓֓֓֓֓֓֓֓֓֓֓֓֓֡֟֜֓֓֓֓֡֡֓֓֡֓֡֓֓֓֡֡֡֡֓֓֡֡֡֡֡֡֡֓֓֡֡֡֡֡֡֡֡	ה מעל	: DVLMTQSPLSLPVTLGQPASISC R	RSSQIIIHSDGNTY-LE		WFQQRPGQSPRLLIY	KVSNRFS	GVPDRFSG
[CDK grafted] KV4B\$HUMAN		DIVMTQSPDSLAVSLGERATINC KSSQSVLYSSNNKNYLA WYQQKPGQPPKLLIY WASTRES	SSQSVLYSSNNK	NYLA WYQ(укрефрекция	WASTRES	GVPDRFSG
(most identical suff N901L/KV4B [Resurfaced]	lins	DVLMTQTPDSLPVSLGDRASISC R	RSSQIIIHSDGNTY-LE WFLQKPGQSPKLLIY KVSNRFS GVPDRFSG	Y-LE WFL(]	QK PGQSPKLLIY	KVSNRFS [L2]	GVPDRFSG
				,			
		0.6 0.8	700	011			
1 N901L		: SGSGTDFTLMISRVEAEDLGVYYC	FQGSHVPHT	FGGGTKLEI	! !	(SEQ ID NO:	NO: 25)
		:SKSGASASLAIGGLQSEDETDYYC AAWDVSLNAYV		FGTGTKVTVL	/L (44)	(SEQ ID	NO: 26)
3 N901L/KOL		SGSGTSFTLAISRVEAEDEGVYYC FQGSHVPHT	FQGSHVPHT	FGGGTKLEI	:- (104)	(SEQ ID	NO: 27)
	(: SGSGTDFTLKISRVEAEDVGVYYC	MQGTHWSWT	FGQGTKVEIK	(87)	(SEQ ID	NO: 28)
car	i has	: SGSGTDFTLKISRVEAEDVGVYYC	FQGSHVPHT	FGGGTKVEI	:- (101)	(SEQ ID	NO: 29)
(CDR glaiced) KV4B\$HUMAN (most identias) curf	1326	: SGSGTDFTLTISSLQAEDVAVYYC	QQYDTIPT 1	FGGGTKVEIK	K (71)	(SEQ ID 1	NO: 30)
(MOSC Idencical S N901L/KV4B [Resurfaced]	i i ine	SGSGTDFTLMISRVEAEDLGVYYC FQGSHVPHT		FGGGTKLEI- J	(109)	(SEQ ID	NO: 31)

FIG. 4B
Heavy Chain Sequences

180	TIY HADTVKG	DQH YADSVKG TIY HADSVKG	NKY YADSVKG TIY YADSVKG	EKY YVDSVKG TIY HADSVKG 	(SEQ ID NO: 32)	(77) (SEQ ID NO: 33)	(106) (SEQ ID NO: 34)	(89) (SEQ ID NO: 35)	(103) (SEQ ID NO: 36)	(74) (SEQ ID NO: 37)	(110) (SEQ ID NO: 38)
170	A YISSGSF	A IIWDDGS A YISSDGF	A VISYDGS	A NIKQDGS A YISSGSF	240 + WGQGTTTVTVS	QGTPVTVS	QGTTVTVS	QGTLVTVS	WGQGTLVTVS		DY WGQGTTVTVS]
160	WVRQAPEKGLEWVA YISSGSFTIY HADTVKG	WVRQAPGKGLEWVA IIWDDGSDQH YADSVKG 	OVQLVESGGGVVQPGRSLRLSCAASGFTFS SYAMH WVRQAPGKGLEWVA VISYDGSNKY YADSVKG: 	:EVQLVESGGGLVQPGGSLRLSCAASGFTFS SYWMS WVRQAPGKGLEWVA NIKQDGSEKY YVDSVKG 	DX	RFTISRNDSKNTLFLOMDSLRPEDTGVYFCAR DGGHGFCSSASCFGPDY WGQGTPVTVS	RFTISRDDPKNTLFLOMTSLRSEDTAMYYCAR MRKGYAMDY WGQGTTVTVS	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR DRKDWGWALFDY WGQGTLVTVS	¥Q		1 -
150	SFGMH	SYAMY SFGMH	SYAMH SFGMH	SYWMS SFGMH	220 +	AR DGGHGI	AR MRKGYA	AR DRKDWC	AR MRKGY?	1R	AR MRKGYA [
140	: DVQLVESGGGLVQPGGSRKLSCAASGFTFS	: EVQLVQSGGGVVQPGRSLRLSCSSSGFIFS 	:QVQLVESGGGVVQPGRSLRLSCAASGFTFS 	URLSCAASGFTFS LRLSCAASGFTFS	190 200 210 230 230)SLRPEDTGVYFC?	rslrsedtamyycz	NSLRAEDTAVYYC	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR MRKGYAM	:RFTISRDNAKNSLYLQMNSLRAEDTAVYYCAR	RFTISRDNAKNTLFLQMTSLRAEDTAMYYCAR MRKGYAM H3
130	SGGGLVQPGGS	SGGGVVQPGRS:	SGGGWQPGRS: 1 SGGGWQPGRS	SGGLVQPGGS	200 + DNPKNTLFLQM	NDSKNTLFLQM	DDPKNTLFLQM	DNSKNTLYLQM	DNSKNTLYLOM	DNAKNSLYLOM	DNAKNTLFLQM
120	: DVQLVE	: EVQLVQ : EVQLVE	OVQLVE:		190 +: :RFTISR	:RFTISR	:RFTISR	:RFTISR	:RFTISR	RFTISR	:RFTISR
	1 N901H	2 KOL 3 N901H/KOL	4 G36005 [most identical seq] 5 N901H/G36005	CDK graited 6 PL0123 Most identical surf 7 N901H/PL0123 Resurfaced	1 N901Н	Z KOL	3 N901H/KOL	4 G36005	(a)	6 PL0123	7 N901H/PL0123 [Resurfaced]

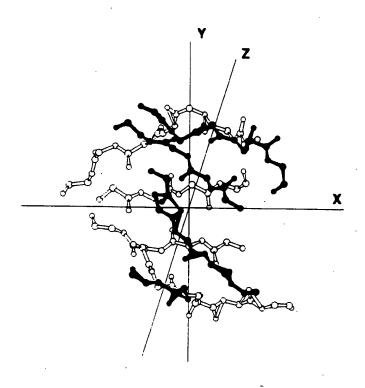
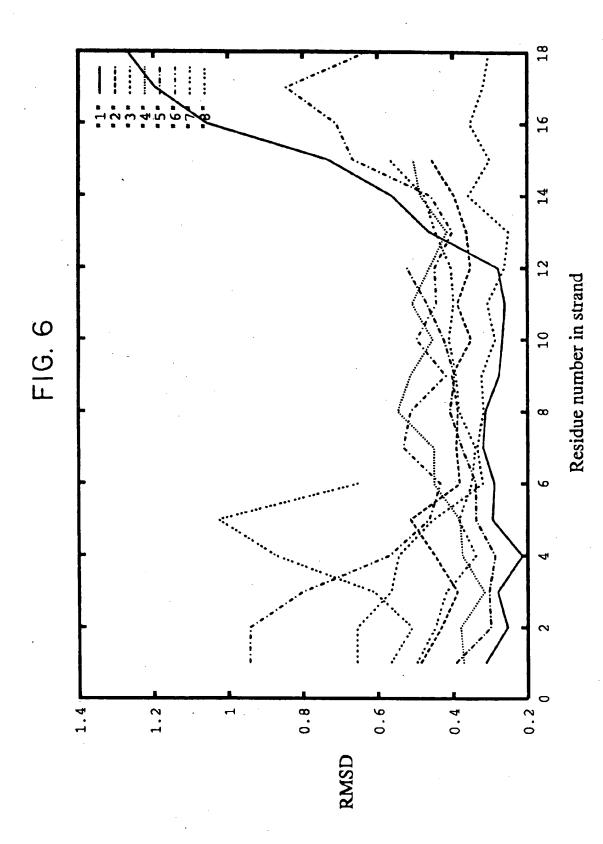
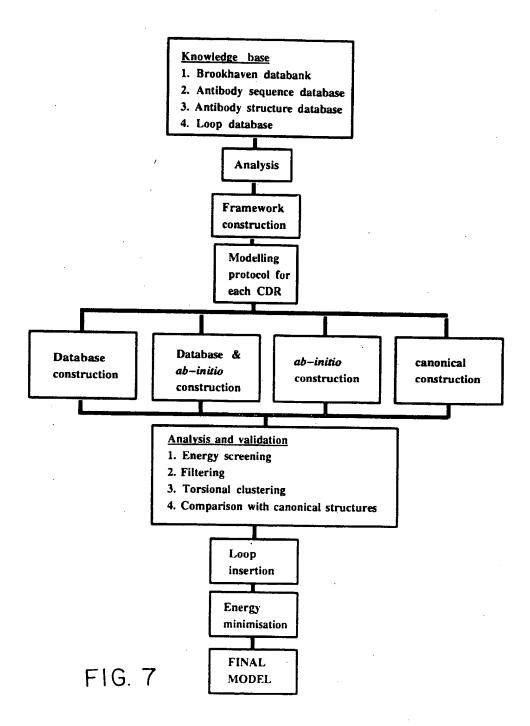
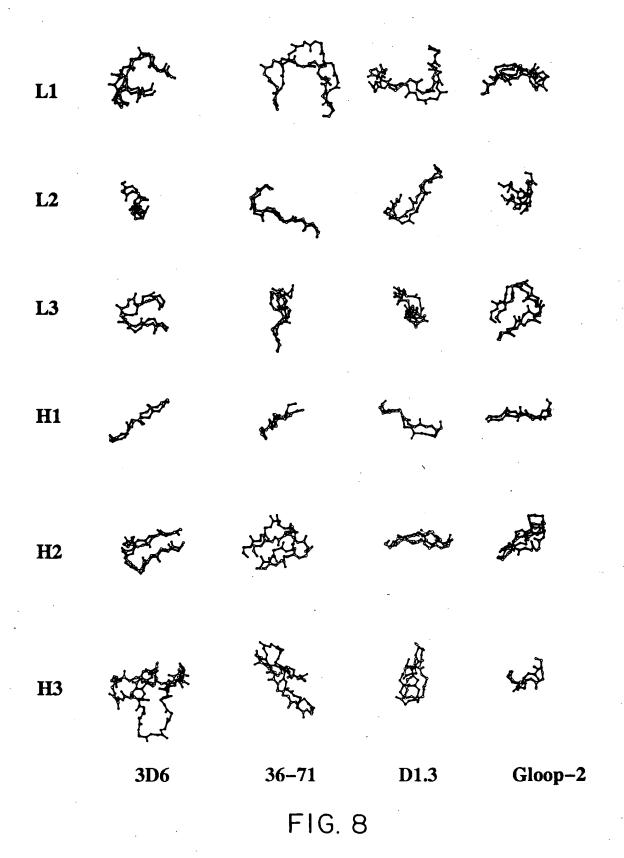
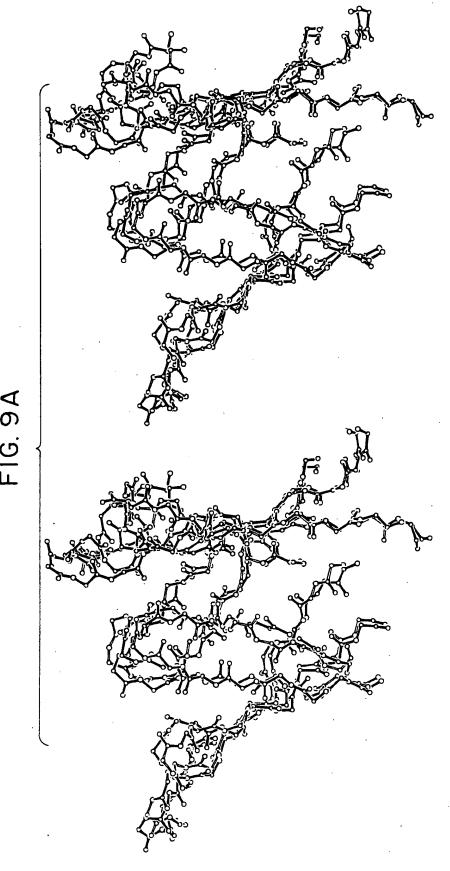


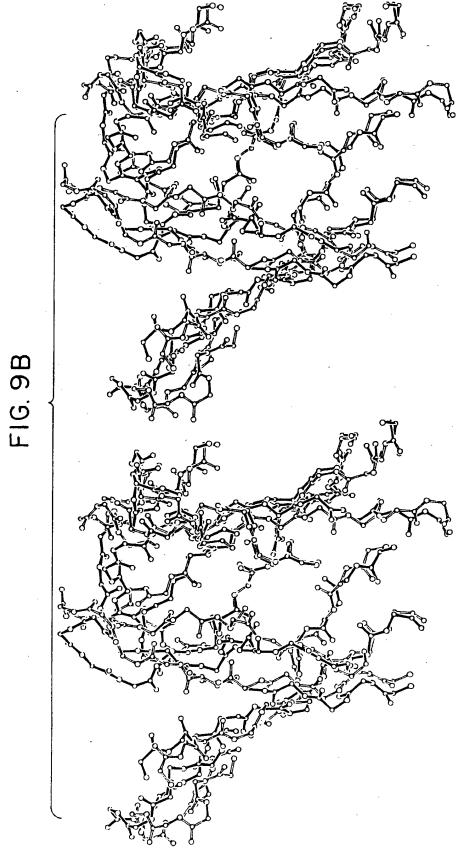
FIG. 5

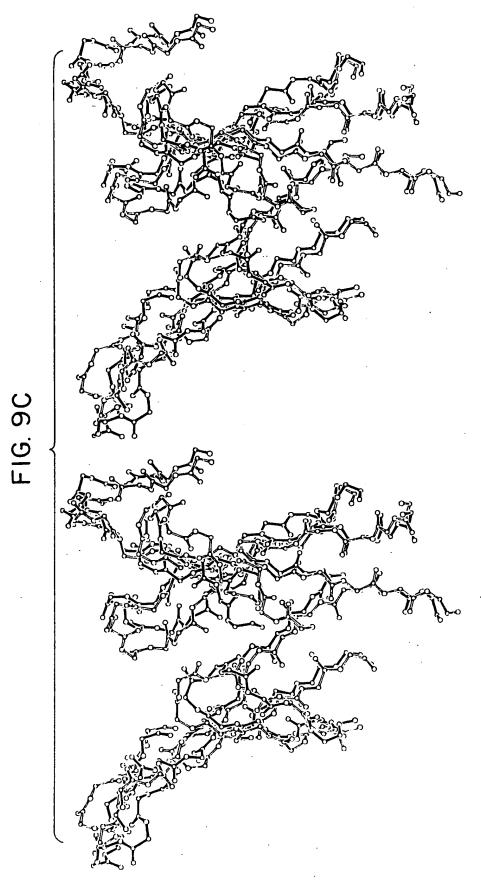


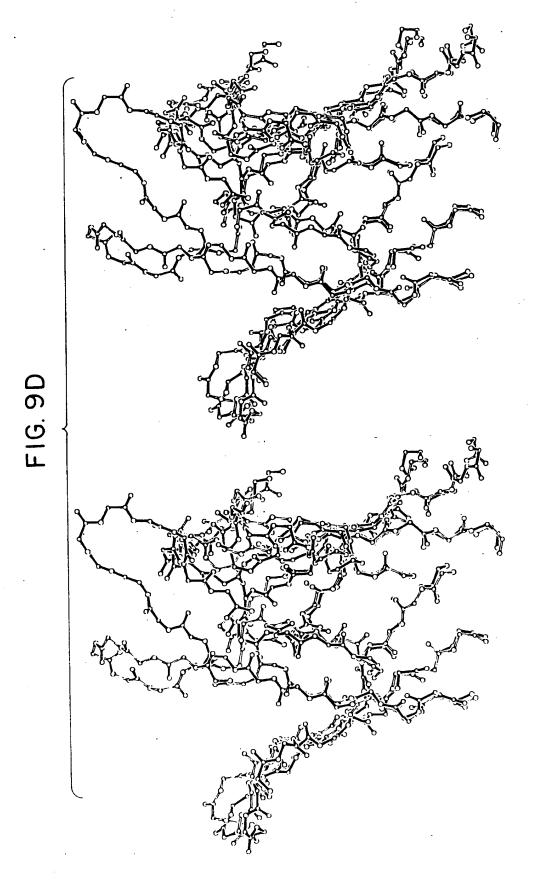












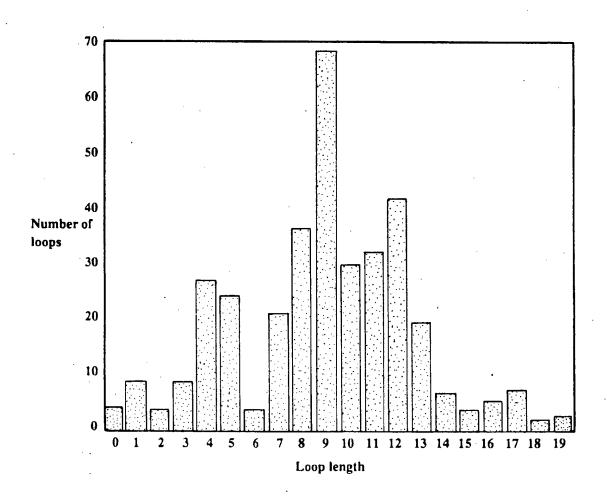


FIG. 10



EUROPEAN SEARCH REPORT

Application Number

EP 93 30 7051

ategory	Citation of document with indica of relevant passage		Relevant to claim	CLASSIFICATION OF THE APPLICATION (Int. Cl.5)
D,A	MOLECULAR IMMUNOLOGY vol. 28, no. 4/5, 1991 pages 489 - 498 PADLAN A E 'POSSIBLE P REDUCING THE IMMUNOGEN VARIABLE DOMAINS WHILE LIGAND-BINDING PROPERT * Materials and Method Tables 1-3 *	ROCEDURE FOR ILCITY OF ANTIBODY PRESERVING THEIR		C12N15/13 C12N15/62 C07K15/00 C12P21/08
D,A	WO-A-9 109 967 (CELLTE 11 July 1991 * p. 5, second paragra paragraph, "Rational"	iph, p. 6 second		
P,A	EP-A-O 519 596 (MERCK 23 December 1992 * Claims *	& CO. INC.)		
				TECHNICAL FIELDS SEARCHED (Int. Cl.5)
				C07K
			,	
				•
			·	
	The present search report has been	drawn up for all claims		
	Place of search	Date of completion of the search		Executer
	MUNICH	12 JANUARY 1994		Germinario C.
Y : pa	CATEGORY OF CITED DOCUMENTS rticularly relevant if taken alone rticularly relevant if combined with another cument of the same category chnological background	E: earlier patent docu after the filing date D: document cited in L: document cited for	ment, but pu the application other reason	blished on, or on